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(54) Title: MOLECULES FOR DISEASE DETECTION AND TREATMENT

(57) Abstract: The present invention provides purified disease detection and treatment molecule polynucleotides (mddt). Also encompassed are the polypeptides (MDDT) encoded by mddt. The invention also provides for the use of mddt, or complements, oligonucleotides, or fragments thereof in diagnostic assays. The invention further provides for vectors and host cells containing mddt for the expression of MDDT. The invention additionally provides for the use of isolated and purified MDDT to induce antibodies and to screen libraries of compounds and the use of anti-MDDT antibodies in diagnostic assays. Also provided are microarrays containing mddt and methods of use.

MOLECULES FOR DISEASE DETECTION AND TREATMENT

TECHNICAL FIELD

The present invention relates to molecules for disease detection and treatment and to the use of these sequences in the diagnosis, study, prevention, and treatment of diseases associated with, as well as effects of exogenous compounds on, the expression of molecules for disease detection and treatment.

BACKGROUND OF THE INVENTION

The human genome is comprised of thousands of genes, many encoding gene products that function in the maintenance and growth of the various cells and tissues in the body. Aberrant expression or mutations in these genes and their products is the cause of, or is associated with, a variety of human diseases such as cancer and other cell proliferative disorders. The identification of these genes and their products is the basis of an ever-expanding effort to find markers for early detection of diseases, and targets for their prevention and treatment.

For example, cancer represents a type of cell proliferative disorder that affects nearly every tissue in the body. A wide variety of molecules, either aberrantly expressed or mutated, can be the cause of, or involved with, various cancers because tissue growth involves complex and ordered 20 patterns of cell proliferation, cell differentiation, and apoptosis. Cell proliferation must be regulated to maintain both the number of cells and their spatial organization. This regulation depends upon the appropriate expression of proteins which control cell cycle progression in response to extracellular signals such as growth factors and other mitogens, and intracellular cues such as DNA damage or nutrient starvation. Molecules which directly or indirectly modulate cell cycle progression fall into 25 several categories, including growth factors and their receptors, second messenger and signal transduction proteins, oncogene products, tumor-suppressor proteins, and mitosis-promoting factors. Aberrant expression or mutations in any of these gene products can result in cell proliferative disorders such as cancer. Oncogenes are genes generally derived from normal genes that, through abnormal expression or mutation, can effect the transformation of a normal cell to a malignant one 30 (oncogenesis). Oncoproteins, encoded by oncogenes, can affect cell proliferation in a variety of ways and include growth factors, growth factor receptors, intracellular signal transducers, nuclear involved in inhibiting cell proliferation. Mutations which cause reduced or loss of function in tumor-suppressor genes result in aberrant cell proliferation and cancer. Thus a wide variety of genes 35 and their products have been found that are associated with cell proliferative disorders such as cancer, but many more may exist that are yet to be discovered.

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DNA-based arrays can provide a simple way to explore the expression of a single polymorphic gene or a large number of genes. When the expression of a single gene is explored, DNA-based arrays are employed to detect the expression of specific gene variants. For example, a p53 tumor suppressor gene array is used to determine whether individuals are carrying mutations that predispose them to cancer. A cytochrome p450 gene array is useful to determine whether individuals have one of a number of specific mutations that could result in increased drug metabolism, drug resistance or drug toxicity.

DNA-based array technology is especially relevant for the rapid screening of expression of a large number of genes. There is a growing awareness that gene expression is affected in a global fashion. A genetic predisposition, disease or therapeutic treatment may affect, directly or indirectly, the expression of a large number of genes. In some cases the interactions may be expected, such as when the genes are part of the same signaling pathway. In other cases, such as when the genes participate in separate signaling pathways, the interactions may be totally unexpected. Therefore, DNA-based arrays can be used to investigate how genetic predisposition, disease, or therapeutic treatment affects the expression of a large number of genes.

The discovery of new molecules for disease detection and treatment satisfies a need in the art by providing new compositions which are useful in the diagnosis, study, prevention, and treatment of diseases associated with, as well as effects of exogenous compounds on, the expression of molecules for disease detection and treatment.

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SUMMARY OF THE INVENTION

The present invention relates to human disease detection and treatment molecule polynucleotides (mddt) as presented in the Sequence Listing. The mddt uniquely identify genes encoding structural, functional, and regulatory disease detection and treatment molecules.

The invention provides an isolated polynucleotide selected from the group consisting of a) a polynucleotide comprising a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; b) a polynucleotide comprising a naturally occurring polynucleotide sequence at least 90% identical to a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; c) a polynucleotide complementary to the polynucleotide of a); d) a polynucleotide complementary to the polynucleotide of b); and e) an RNA equivalent of a) through d). In one alternative, the polynucleotide comprises a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104. In another alternative, the polynucleotide comprises at least 30 contiguous nucleotides of a polynucleotide sequence selected from the group consisting of a) a polynucleotide comprising a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; b) a polynucleotide comprising a naturally occurring polynucleotide comprising a polynucleotide

sequence at least 90% identical to a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; c) a polynucleotide complementary to the polynucleotide of a); d) a polynucleotide complementary to the polynucleotide of b); and e) an RNA equivalent of a) through d). In another alternative, the polynucleotide comprises at least 60 contiguous nucleotides of a 5 polynucleotide selected from the group consisting of a) a polynucleotide comprising a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; b) a polynucleotide comprising a naturally occurring polynucleotide comprising a polynucleotide sequence at least 90% identical to a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; c) a polynucleotide complementary to the polynucleotide of a); d) a polynucleotide complementary to the 10 polynucleotide of b); and e) an RNA equivalent of a) through d). The invention further provides a composition for the detection of expression of disease detection and treatment molecule polynucleotides comprising at least one isolated polynucleotide comprising a polynucleotide selected from the group consisting of a) a polynucleotide comprising a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; b) a polynucleotide comprising a naturally occurring polynucleotide sequence at least 90% identical to a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; c) a polynucleotide complementary to the polynucleotide of a); d) a polynucleotide complementary to the polynucleotide of b); and e) an RNA equivalent of a) through d); and a detectable label.

target polynucleotide comprising a polynucleotide sequence of a polynucleotide selected from the group consisting of a) a polynucleotide comprising a polynucleotide sequence of a polynucleotide selected from the group consisting of SEQ ID NO:1-104; b) a polynucleotide comprising a naturally occurring polynucleotide sequence at least 90% identical to a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; c) a polynucleotide complementary to the polynucleotide of a); d) a polynucleotide complementary to the polynucleotide of a) through d). The method comprises a) amplifying said target polynucleotide or fragment thereof using polymerase chain reaction amplification, and b) detecting the presence or absence of said amplified target polynucleotide or fragment thereof, and, optionally, if present, the amount thereof.

The invention also provides a method for detecting a target polynucleotide in a sample, said target polynucleotide having a polynucleotide selected from the group consisting of a) a polynucleotide comprising a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; b) a polynucleotide comprising a naturally occurring polynucleotide sequence at least 90% identical to a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; c) a polynucleotide complementary to the polynucleotide of a); d) a polynucleotide complementary to the polynucleotide of b); and e) an RNA equivalent of a) through d). The method comprises a)

hybridizing the sample with a probe comprising at least 20 contiguous nucleotides comprising a sequence complementary to said target polynucleotide in the sample, and which probe specifically hybridizes to said target polynucleotide, under conditions whereby a hybridization complex is formed between said probe and said target polynucleotide, and b) detecting the presence or absence of said 5 hybridization complex, and, optionally, if present, the amount thereof. In one alternative, the invention provides a composition comprising a target polynucleotide of the method, wherein said probe comprises at least 30 contiguous nucleotides. In one alternative, the invention provides a composition comprising a target polynucleotide of the method, wherein said probe comprises at least 60 contiguous nucleotides.

The invention further provides a recombinant polynucleotide comprising a promoter sequence operably linked to an isolated polynucleotide selected from the group consisting of a) a polynucleotide comprising a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; b) a polynucleotide comprising a naturally occurring polynucleotide sequence at least 90% identical to a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; 15 c) a polynucleotide complementary to the polynucleotide of a); d) a polynucleotide complementary to the polynucleotide of b); and e) an RNA equivalent of a) through d). In one alternative, the invention provides a cell transformed with the recombinant polynucleotide. In another alternative, the invention provides a transgenic organism comprising the recombinant polynucleotide.

The invention also provides a method for producing a disease detection and treatment 20 molecule polypeptide, the method comprising a) culturing a cell under conditions suitable for expression of the disease detection and treatment molecule polypeptide, wherein said cell is transformed with a recombinant polynucleotide, said recombinant polynucleotide comprising an isolated polynucleotide selected from the group consisting of i) a polynucleotide comprising a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; ii) a 25 polynucleotide comprising a naturally occurring polynucleotide sequence at least 90% identical to a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; iii) a polynucleotide complementary to the polynucleotide of i); iv) a polynucleotide complementary to the polynucleotide of ii); and v) an RNA equivalent of i) through iv), and b) recovering the disease detection and treatment molecule polypeptide so expressed. The invention additionally provides a 30 method wherein the polypeptide has an amino acid sequence selected from the group consisting of SEQ ID NO:105-208.

The invention also provides an isolated disease detection and treatment molecule polypeptide (MDDT) encoded by at least one polynucleotide comprising a polynucleotide sequence selected from the group consisting of SEO ID NO:1-104. The invention further provides a method of screening for 35 a test compound that specifically binds to the polypeptide having an amino acid sequence selected

from the group consisting of SEQ ID NO:105-208. The method comprises a) combining the polypeptide having an amino acid sequence selected from the group consisting of SEO ID NO:105-208 with at least one test compound under suitable conditions, and b) detecting binding of the polypeptide having an amino acid sequence selected from the group consisting of SEO ID NO:105-5 208 to the test compound, thereby identifying a compound that specifically binds to the polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:105-208.

The invention further provides a microarray wherein at least one element of the microarray is an isolated polynucleotide comprising at least 30 contiguous nucleotides of a polynucleotide selected from the group consisting of a) a polynucleotide comprising a polynucleotide sequence selected from 10 the group consisting of SEQ ID NO:1-104; b) a polynucleotide comprising a naturally occurring polynucleotide sequence at least 90% identical to a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; c) a polynucleotide complementary to the polynucleotide of a); d) a polynucleotide complementary to the polynucleotide of b); and e) an RNA equivalent of a) through d). The invention also provides a method for generating a transcript image of a sample which 15 contains polynucleotides. The method comprises a) labeling the polynucleotides of the sample, b) contacting the elements of the microarray with the labeled polynucleotides of the sample under conditions suitable for the formation of a hybridization complex, and c) quantifying the expression of the polynucleotides in the sample.

Additionally, the invention provides a method for screening a compound for effectiveness in 20 altering expression of a target polynucleotide, wherein said target polynucleotide comprises a polynucleotide selected from the group consisting of a) a polynucleotide comprising a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; b) a polynucleotide comprising a naturally occurring polynucleotide sequence at least 90% identical to a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; c) a polynucleotide complementary to the 25 polynucleotide of a); d) a polynucleotide complementary to the polynucleotide of b); and e) an RNA equivalent of a) through d). The method comprises a) exposing a sample comprising the target polynucleotide to a compound, b) detecting altered expression of the target polynucleotide, and c) comparing the expression of the target polynucleotide in the presence of varying amounts of the compound and in the absence of the compound.

The invention further provides a method for assessing toxicity of a test compound, said method comprising a) treating a biological sample containing nucleic acids with the test compound; b) hybridizing the nucleic acids of the treated biological sample with a probe comprising at least 20 contiguous nucleotides of a polynucleotide selected from the group consisting of i) a polynucleotide comprising a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; ii) a 35 polynucleotide comprising a naturally occurring polynucleotide sequence at least 90% identical to a

polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; iii) a polynucleotide complementary to the polynucleotide of i); iv) a polynucleotide complementary to the polynucleotide of ii); and v) an RNA equivalent of i) through iv). Hybridization occurs under conditions whereby a specific hybridization complex is formed between said probe and a target 5 polynucleotide in the biological sample, said target polynucleotide comprising a polynucleotide sequence of a polynucleotide selected from the group consisting of i) a polynucleotide comprising a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; ii) a polynucleotide comprising a naturally occurring polynucleotide sequence at least 90% identical to a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104; iii) a 10 polynucleotide complementary to the polynucleotide of i); iv) a polynucleotide complementary to the polynucleotide of ii); and v) an RNA equivalent of i) through iv), and alternatively, the target polynucleotide comprises a polynucleotide sequence of a fragment of a polynucleotide selected from the group consisting of i-v above; c) quantifying the amount of hybridization complex; and d) comparing the amount of hybridization complex in the treated biological sample with the amount of 15 hybridization complex in an untreated biological sample, wherein a difference in the amount of hybridization complex in the treated biological sample is indicative of toxicity of the test compound.

The invention further provides an isolated polypeptide selected from the group consisting of a) a polypeptide comprising an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, b) a polypeptide comprising a naturally occurring amino acid sequence at least 90% identical to an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, c) a biologically active fragment of a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, and d) an immunogenic fragment of a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:105-208. In one alternative, the invention provides an isolated polypeptide comprising an amino acid sequence selected from the group consisting of SEQ ID NO:105-208.

The invention further provides an isolated polynucleotide encoding a polypeptide selected from the group consisting of a) a polypeptide comprising an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, b) a polypeptide comprising a naturally occurring amino acid sequence at least 90% identical to an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, c) a biologically active fragment of a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, and d) an immunogenic fragment of a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:105-208. In one alternative, the polynucleotide encodes a polypeptide comprising an amino acid sequence selected from the group consisting of SEQ ID NO:105-208. In another alternative, the polynucleotide comprises a polynucleotide sequence selected from the group consisting of SEQ ID NO:105-208.

NO:1-104.

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Additionally, the invention provides an isolated antibody which specifically binds to a polypeptide selected from the group consisting of a) a polypeptide comprising an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, b) a polypeptide comprising a 5 naturally occurring amino acid sequence at least 90% identical to an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, c) a biologically active fragment of a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, and d) an immunogenic fragment of a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:105-208.

The invention further provides a composition comprising a polypeptide selected from the group consisting of a) a polypeptide comprising an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, b) a polypeptide comprising a naturally occurring amino acid sequence at least 90% identical to an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, c) a biologically active fragment of a polypeptide having an amino acid sequence 15 selected from the group consisting of SEQ ID NO:105-208, and d) an immunogenic fragment of a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, and a pharmaceutically acceptable excipient. In one embodiment, the composition comprises a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:105-208. The invention additionally provides a method of treating a disease or condition associated with 20 decreased expression of functional MDDT, comprising administering to a patient in need of such treatment the composition.

The invention also provides a method for screening a compound for effectiveness as an agonist of a polypeptide selected from the group consisting of a) a polypeptide comprising an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, b) a polypeptide 25 comprising a naturally occurring amino acid sequence at least 90% identical to an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, c) a biologically active fragment of a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, and d) an immunogenic fragment of a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:105-208. The method comprises a) exposing a 30 sample comprising the polypeptide to a compound, and b) detecting agonist activity in the sample. In one alternative, the invention provides a composition comprising an agonist compound identified by the method and a pharmaceutically acceptable excipient. In another alternative, the invention provides a method of treating a disease or condition associated with decreased expression of functional MDDT, comprising administering to a patient in need of such treatment the composition.

Additionally, the invention provides a method for screening a compound for effectiveness as

an antagonist of a polypeptide selected from the group consisting of a) a polypeptide comprising an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, b) a polypeptide comprising a naturally occurring amino acid sequence at least 90% identical to an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, c) a biologically active

5 fragment of a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, and d) an immunogenic fragment of a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:105-208. The method comprises a) exposing a sample comprising the polypeptide to a compound, and b) detecting antagonist activity in the sample. In one alternative, the invention provides a composition comprising an antagonist compound

10 identified by the method and a pharmaceutically acceptable excipient. In another alternative, the invention provides a method of treating a disease or condition associated with overexpression of functional MDDT, comprising administering to a patient in need of such treatment the composition.

The invention further provides a method of screening for a compound that modulates the activity of a polypeptide selected from the group consisting of a) a polypeptide comprising an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, b) a polypeptide comprising a naturally occurring amino acid sequence at least 90% identical to an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, c) a biologically active fragment of a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, and d) an immunogenic fragment of a polypeptide having an amino acid sequence selected from the group consisting of SEQ ID NO:105-208. The method comprises a) combining the polypeptide with at least one test compound under conditions permissive for the activity of the polypeptide, b) assessing the activity of the polypeptide in the presence of the test compound, and c) comparing the activity of the polypeptide in the presence of the test compound with the activity of the polypeptide in the presence of the test compound with the activity of the polypeptide in the presence of the test compound that modulates the activity of the polypeptide.

DESCRIPTION OF THE TABLES

Table 1 shows the sequence identification numbers (SEQ ID NO:s) and template

identification numbers (template IDs) corresponding to the polynucleotides of the present invention,
along with the sequence identification numbers (SEQ ID NO:s) and open reading frame identification
numbers (ORF IDs) corresponding to polypeptides encoded by the template ID.

Table 2 shows the sequence identification numbers (SEQ ID NO:s) and template identification numbers (template IDs) corresponding to the polynucleotides of the present invention, along with their GenBank hits (GI Numbers), probability scores, and functional annotations

corresponding to the GenBank hits.

Table 3 shows the sequence identification numbers (SEQ ID NO:s) and template identification numbers (template IDs) corresponding to the polynucleotides of the present invention, along with polynucleotide segments of each template sequence as defined by the indicated "start" and "stop" nucleotide positions. The reading frames of the polynucleotide segments and the Pfam hits, Pfam descriptions, and E-values corresponding to the polypeptide domains encoded by the polynucleotide segments are indicated.

Table 4 shows the sequence identification numbers (SEQ ID NO:s) and template identification numbers (template IDs) corresponding to the polynucleotides of the present invention, along with polynucleotide segments of each template sequence as defined by the indicated "start" and "stop" nucleotide positions. The reading frames of the polynucleotide segments are shown, and the polypeptides encoded by the polynucleotide segments constitute either signal peptide (SP) or transmembrane (TM) domains, as indicated. For TM domains, the membrane topology of the encoded polypeptide sequence is indicated as being transmembrane or on the cytosolic or non-

Table 5 shows the sequence identification numbers (SEQ ID NO:s) and template identification numbers (template IDs) corresponding to the polynucleotides of the present invention, along with the component sequence identification spans (component spans) corresponding to each template. The component sequences, which were used to assemble the template sequences, are defined by the spans indicating the nucleotide positions along each template.

Table 6 shows the tissue distribution profiles for the templates of the invention.

Table 7 shows the sequence identification numbers (SEQ ID NO:s) corresponding to the polypeptides of the present invention, along with the reading frames used to obtain the polypeptide segments, the lengths of the polypeptide segments, the "start" and "stop" nucleotide positions of the polypeptide sequences used to define the encoded polypeptide segments, the GenBank hits (GI Numbers), probability scores, and functional annotations corresponding to the GenBank hits.

Table 8 summarizes the bioinformatics tools which are useful for analysis of the polynucleotides of the present invention. The first column of Table 8 lists analytical tools, programs, and algorithms, the second column provides brief descriptions thereof, the third column presents appropriate references, all of which are incorporated by reference herein in their entirety, and the fourth column presents, where applicable, the scores, probability values, and other parameters used to evaluate the strength of a match between two sequences (the higher the score, the greater the homology between two sequences).

DETAILED DESCRIPTION OF THE INVENTION

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Before the nucleic acid sequences and methods are presented, it is to be understood that this invention is not limited to the particular machines, methods, and materials described. Although particular embodiments are described, machines, methods, and materials similar or equivalent to these embodiments may be used to practice the invention. The preferred machines, methods, and materials set forth are not intended to limit the scope of the invention which is limited only by the appended claims.

The singular forms "a", "an", and "the" include plural reference unless the context clearly dictates otherwise. All technical and scientific terms have the meanings commonly understood by one of ordinary skill in the art. All publications are incorporated by reference for the purpose of describing and disclosing the cell lines, vectors, and methodologies which are presented and which might be used in connection with the invention. Nothing in the specification is to be construed as an admission that the invention is not entitled to antedate such disclosure by virtue of prior invention.

Definitions

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As used herein, the lower case "mddt" refers to a nucleic acid sequence, while the upper case "MDDT" refers to an amino acid sequence encoded by mddt. A "full-length" mddt refers to a nucleic acid sequence containing the entire coding region of a gene endogenously expressed in human tissue.

"Adjuvants" are materials such as Freund's adjuvant, mineral gels (aluminum hydroxide), and surface active substances (lysolecithin, pluronic polyols, polyanions, peptides, oil emulsions, keyhole limpet hemocyanin, and dinitrophenol) which may be administered to increase a host's immunological response.

"Allele" refers to an alternative form of a nucleic acid sequence. Alleles result from a "mutation," a change or an alternative reading of the genetic code. Any given gene may have none, one, or many allelic forms. Mutations which give rise to alleles include deletions, additions, or substitutions of nucleotides. Each of these changes may occur alone, or in combination with the others, one or more times in a given nucleic acid sequence. The present invention encompasses allelic mddt.

An "allelic variant" is an alternative form of the gene encoding MDDT. Allelic variants may result from at least one mutation in the nucleic acid sequence and may result in altered mRNAs or in polypeptides whose structure or function may or may not be altered. A gene may have none, one, or many allelic variants of its naturally occurring form. Common mutational changes which give rise to allelic variants are generally ascribed to natural deletions, additions, or substitutions of

nucleotides. Each of these types of changes may occur alone, or in combination with the others, one or more times in a given sequence.

"Altered" nucleic acid sequences encoding MDDT include those sequences with deletions, insertions, or substitutions of different nucleotides, resulting in a polypeptide the same as MDDT or a polypeptide with at least one functional characteristic of MDDT. Included within this definition are polymorphisms which may or may not be readily detectable using a particular oligonucleotide probe of the polynucleotide encoding MDDT, and improper or unexpected hybridization to allelic variants, with a locus other than the normal chromosomal locus for the polynucleotide sequence encoding MDDT. The encoded protein may also be "altered," and may contain deletions, insertions, or substitutions of amino acid residues which produce a silent change and result in a functionally equivalent MDDT. Deliberate amino acid substitutions may be made on the basis of similarity in polarity, charge, solubility, hydrophobicity, hydrophilicity, and/or the amphipathic nature of the residues, as long as the biological or immunological activity of MDDT is retained. For example, negatively charged amino acids may include aspartic acid and glutamic acid, and positively charged amino acids may include lysine and arginine. Amino acids with uncharged polar side chains having similar hydrophilicity values may include: asparagine and glutamine; and serine and threonine. Amino acids with uncharged side chains having similar hydrophilicity values may include: leucine, isoleucine, and valine; glycine and alanine; and phenylalanine and tyrosine.

"Amino acid sequence" refers to a peptide, a polypeptide, or a protein of either natural or synthetic origin. The amino acid sequence is not limited to the complete, endogenous amino acid sequence and may be a fragment, epitope, variant, or derivative of a protein expressed by a nucleic acid sequence.

"Amplification" refers to the production of additional copies of a sequence and is carried out using polymerase chain reaction (PCR) technologies well known in the art.

"Antibody" refers to intact molecules as well as to fragments thereof, such as Fab, F(ab')₂, and Fv fragments, which are capable of binding the epitopic determinant. Antibodies that bind MDDT polypeptides can be prepared using intact polypeptides or using fragments containing small peptides of interest as the immunizing antigen. The polypeptide or peptide used to immunize an animal (e.g., a mouse, a rat, or a rabbit) can be derived from the translation of RNA, or synthesized chemically, and can be conjugated to a carrier protein if desired. Commonly used carriers that are chemically coupled to peptides include bovine serum albumin, thyroglobulin, and keyhole limpet hemocyanin (KLH). The coupled peptide is then used to immunize the animal.

The term "aptamer" refers to a nucleic acid or oligonucleotide molecule that binds to a specific molecular target. Aptamers are derived from an <u>in vitro</u> evolutionary process (e.g., SELEX (Systematic Evolution of Ligands by EXponential Enrichment), described in U.S. Patent No.

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5,270,163), which selects for target-specific aptamer sequences from large combinatorial libraries. Aptamer compositions may be double-stranded or single-stranded, and may include deoxyribonucleotides, ribonucleotides, nucleotide derivatives, or other nucleotide-like molecules. The nucleotide components of an aptamer may have modified sugar groups (e.g., the 2'-OH group of a ribonucleotide may be replaced by 2'-F or 2'-NH₂), which may improve a desired property, e.g., resistance to nucleases or longer lifetime in blood. Aptamers may be conjugated to other molecules, e.g., a high molecular weight carrier to slow clearance of the aptamer from the circulatory system. Aptamers may be specifically cross-linked to their cognate ligands, e.g., by photo-activation of a cross-linker. (See, e.g., Brody, E.N. and L. Gold (2000) J. Biotechnol. 74:5-13.)

The term "intramer" refers to an aptamer which is expressed <u>in vivo</u>. For example, a vaccinia virus-based RNA expression system has been used to express specific RNA aptamers at high levels in the cytoplasm of leukocytes (Blind, M. et al. (1999) Proc. Natl Acad. Sci. USA 96:3606-3610).

The term "spiegelmer" refers to an aptamer which includes L-DNA, L-RNA, or other left-handed nucleotide derivatives or nucleotide-like molecules. Aptamers containing left-handed nucleotides are resistant to degradation by naturally occurring enzymes, which normally act on substrates containing right-handed nucleotides.

"Antisense sequence" refers to a sequence capable of specifically hybridizing to a target sequence. The antisense sequence may include DNA, RNA, or any nucleic acid mimic or analog such as peptide nucleic acid (PNA); oligonucleotides having modified backbone linkages such as phosphorothioates, methylphosphonates, or benzylphosphonates; oligonucleotides having modified sugar groups such as 2'-methoxyethyl sugars or 2'-methoxyethoxy sugars; or oligonucleotides having modified bases such as 5-methyl cytosine, 2'-deoxyuracil, or 7-deaza-2'-deoxyguanosine.

"Antisense technology" refers to any technology which relies on the specific hybridization of an antisense sequence to a target sequence.

A "bin" is a portion of computer memory space used by a computer program for storage of data, and bounded in such a manner that data stored in a bin may be retrieved by the program.

"Biologically active" refers to an amino acid sequence having a structural, regulatory, or biochemical function of a naturally occurring amino acid sequence.

"Clone joining" is a process for combining gene bins based upon the bins' containing sequence information from the same clone. The sequences may assemble into a primary gene transcript as well as one or more splice variants.

"Complementary" describes the relationship between two single-stranded nucleic acid sequences that anneal by base-pairing (5'-A-G-T-3' pairs with its complement 3'-T-C-A-5').

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A "component sequence" is a nucleic acid sequence selected by a computer program such as PHRED and used to assemble a consensus or template sequence from one or more component sequences.

A "consensus sequence" or "template sequence" is a nucleic acid sequence which has been assembled from overlapping sequences, using a computer program for fragment assembly such as the GELVIEW fragment assembly system (Genetics Computer Group (GCG), Madison WI) or using a relational database management system (RDMS).

"Conservative amino acid substitutions" are those substitutions that, when made, least interfere with the properties of the original protein, i.e., the structure and especially the function of the protein is conserved and not significantly changed by such substitutions. The table below shows amino acids which may be substituted for an original amino acid in a protein and which are regarded as conservative substitutions.

	Original Residue	Conservative Substitution
15	Ala	Gly, Ser
	Arg	His, Lys
	Asn	Asp, Gln, His
	Asp	Asn, Glu
	Cys	Ala, Ser
20	Gln	Asn, Glu, His
	Glu	Asp, Gln, His
	Gly	Ala
	His	Asn, Arg, Gln, Glu
	Ile	Leu, Val
25 .	Leu	Ile, Val
	Lys	Arg, Gln, Glu
	Met	Leu, Ile
	Phe	His, Met, Leu, Trp, Tyr
	Ser	Cys, Thr
30	Thr	Ser, Val
	Тгр	Phe, Tyr
	Tyr	His, Phe, Trp
	Val	Ile, Leu, Thr

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Conservative substitutions generally maintain (a) the structure of the polypeptide backbone in the area of the substitution, for example, as a beta sheet or alpha helical conformation, (b) the charge or hydrophobicity of the molecule at the target site, or (c) the bulk of the side chain.

"Deletion" refers to a change in either a nucleic or amino acid sequence in which at least one nucleotide or amino acid residue, respectively, is absent.

"Derivative" refers to the chemical modification of a nucleic acid sequence, such as by replacement of hydrogen by an alkyl, acyl, amino, hydroxyl, or other group.

"Differential expression" refers to increased or upregulated; or decreased, downregulated, or absent gene or protein expression, determined by comparing at least two different samples. Such comparisons may be carried out between, for example, a treated and an untreated sample, or a diseased and a normal sample.

The terms "element" and "array element" refer to a polynucleotide, polypeptide, or other chemical compound having a unique and defined position on a microarray.

The term "modulate" refers to a change in the activity of MDDT. For example, modulation may cause an increase or a decrease in protein activity, binding characteristics, or any other biological, functional, or immunological properties of MDDT.

"E-value" refers to the statistical probability that a match between two sequences occurred by chance.

"Exon shuffling" refers to the recombination of different coding regions (exons). Since an exon may represent a structural or functional domain of the encoded protein, new proteins may be assembled through the novel reassortment of stable substructures, thus allowing acceleration of the evolution of new protein functions.

A "fragment" is a unique portion of mddt or MDDT which is identical in sequence to but shorter in length than the parent sequence. A fragment may comprise up to the entire length of the defined sequence, minus one nucleotide/amino acid residue. For example, a fragment may comprise from 10 to 1000 contiguous amino acid residues or nucleotides. A fragment used as a probe, primer, antigen, therapeutic molecule, or for other purposes, may be at least 5, 10, 15, 16, 20, 25, 30, 40, 50, 60, 75, 100, 150, 250 or at least 500 contiguous amino acid residues or nucleotides in length. Fragments may be preferentially selected from certain regions of a molecule. For example, a polypeptide fragment may comprise a certain length of contiguous amino acids selected from the first 250 or 500 amino acids (or first 25% or 50%) of a polypeptide as shown in a certain defined sequence. Clearly these lengths are exemplary, and any length that is supported by the specification, including the Sequence Listing and the figures, may be encompassed by the present embodiments.

A fragment of mddt comprises a region of unique polynucleotide sequence that specifically identifies mddt, for example, as distinct from any other sequence in the same genome. A fragment of mddt is useful, for example, in hybridization and amplification technologies and in analogous methods that distinguish mddt from related polynucleotide sequences. The precise length of a fragment of mddt and the region of mddt to which the fragment corresponds are routinely determinable by one of ordinary skill in the art based on the intended purpose for the fragment.

A fragment of MDDT is encoded by a fragment of mddt. A fragment of MDDT comprises a region of unique amino acid sequence that specifically identifies MDDT. For example, a fragment of MDDT is useful as an immunogenic peptide for the development of antibodies that specifically

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recognize MDDT. The precise length of a fragment of MDDT and the region of MDDT to which the fragment corresponds are routinely determinable by one of ordinary skill in the art based on the intended purpose for the fragment.

A "full length" nucleotide sequence is one containing at least a start site for translation to a protein sequence, followed by an open reading frame and a stop site, and encoding a "full length" polypeptide.

"Hit" refers to a sequence whose annotation will be used to describe a given template. Criteria for selecting the top hit are as follows: if the template has one or more exact nucleic acid matches, the top hit is the exact match with highest percent identity. If the template has no exact matches but has significant protein hits, the top hit is the protein hit with the lowest E-value. If the template has no significant protein hits, but does have significant non-exact nucleotide hits, the top hit is the nucleotide hit with the lowest E-value.

"Homology" refers to sequence similarity either between a reference nucleic acid sequence and at least a fragment of an mddt or between a reference amino acid sequence and a fragment of an MDDT.

"Hybridization" refers to the process by which a strand of nucleotides anneals with a complementary strand through base pairing. Specific hybridization is an indication that two nucleic acid sequences share a high degree of identity. Specific hybridization complexes form under defined annealing conditions, and remain hybridized after the "washing" step. The defined hybridization conditions include the annealing conditions and the washing step(s), the latter of which is particularly important in determining the stringency of the hybridization process, with more stringent conditions allowing less non-specific binding, i.e., binding between pairs of nucleic acid probes that are not perfectly matched. Permissive conditions for annealing of nucleic acid sequences are routinely determinable and may be consistent among hybridization experiments, whereas wash conditions may be varied among experiments to achieve the desired stringency.

Generally, stringency of hybridization is expressed with reference to the temperature under which the wash step is carried out. Generally, such wash temperatures are selected to be about 5°C to 20°C lower than the thermal melting point (T_m) for the specific sequence at a defined ionic strength and pH. The T_m is the temperature (under defined ionic strength and pH) at which 50% of the target sequence hybridizes to a perfectly matched probe. An equation for calculating T_m and conditions for nucleic acid hybridization is well known and can be found in Sambrook et al., 1989, Molecular Cloning: A Laboratory Manual, 2^{nd} ed., vol. 1-3, Cold Spring Harbor Press, Plainview NY; specifically see volume 2, chapter 9.

High stringency conditions for hybridization between polynucleotides of the present invention include wash conditions of 68°C in the presence of about 0.2 x SSC and about 0.1% SDS,

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for 1 hour. Alternatively, temperatures of about 65°C, 60°C, or 55°C may be used. SSC concentration may be varied from about 0.2 to 2 x SSC, with SDS being present at about 0.1%. Typically, blocking reagents are used to block non-specific hybridization. Such blocking reagents include, for instance, denatured salmon sperm DNA at about 100-200 μ g/ml. Useful variations on these conditions will be readily apparent to those skilled in the art. Hybridization, particularly under high stringency conditions, may be suggestive of evolutionary similarity between the nucleotides. Such similarity is strongly indicative of a similar role for the nucleotides and their resultant proteins.

Other parameters, such as temperature, salt concentration, and detergent concentration may be varied to achieve the desired stringency. Denaturants, such as formamide at a concentration of about 35-50% v/v, may also be used under particular circumstances, such as RNA:DNA hybridizations. Appropriate hybridization conditions are routinely determinable by one of ordinary skill in the art.

"Immunologically active" or "immunogenic" describes the potential for a natural, recombinant, or synthetic peptide, epitope, polypeptide, or protein to induce antibody production in appropriate animals, cells, or cell lines.

"Immune response" can refer to conditions associated with inflammation, trauma, immune disorders, or infectious or genetic disease, etc. These conditions can be characterized by expression of various factors, e.g., cytokines, chemokines, and other signaling molecules, which may affect cellular and systemic defense systems.

An "immunogenic fragment" is a polypeptide or oligopeptide fragment of MDDT which is capable of eliciting an immune response when introduced into a living organism, for example, a mammal. The term "immunogenic fragment" also includes any polypeptide or oligopeptide fragment of MDDT which can be useful in any of the antibody production methods disclosed herein or known in the art.

"Insertion" or "addition" refers to a change in either a nucleic or amino acid sequence in which at least one nucleotide or residue, respectively, is added to the sequence.

"Labeling" refers to the covalent or noncovalent joining of a polynucleotide, polypeptide, or antibody with a reporter molecule capable of producing a detectable or measurable signal.

"Microarray" is any arrangement of nucleic acids, amino acids, antibodies, etc., on a substrate. The substrate may be a solid support such as beads, glass, paper, nitrocellulose, nylon, or an appropriate membrane.

"Linkers" are short stretches of nucleotide sequence which may be added to a vector or an mddt to create restriction endonuclease sites to facilitate cloning. "Polylinkers" are engineered to incorporate multiple restriction enzyme sites and to provide for the use of enzymes which leave 5' or 3' overhangs (e.g., BamHI, EcoRI, and HindIII) and those which provide blunt ends (e.g., EcoRV,

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SnaBI, and StuI).

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"Naturally occurring" refers to an endogenous polynucleotide or polypeptide that may be isolated from viruses or prokaryotic or eukaryotic cells.

"Nucleic acid sequence" refers to the specific order of nucleotides joined by phosphodiester bonds in a linear, polymeric arrangement. Depending on the number of nucleotides, the nucleic acid sequence can be considered an oligomer, oligonucleotide, or polynucleotide. The nucleic acid can be DNA, RNA, or any nucleic acid analog, such as PNA, may be of genomic or synthetic origin, may be either double-stranded or single-stranded, and can represent either the sense or antisense (complementary) strand.

"Oligomer" refers to a nucleic acid sequence of at least about 6 nucleotides and as many as about 60 nucleotides, preferably about 15 to 40 nucleotides, and most preferably between about 20 and 30 nucleotides, that may be used in hybridization or amplification technologies. Oligomers may be used as, e.g., primers for PCR, and are usually chemically synthesized.

"Operably linked" refers to the situation in which a first nucleic acid sequence is placed in a functional relationship with the second nucleic acid sequence. For instance, a promoter is operably linked to a coding sequence if the promoter affects the transcription or expression of the coding sequence. Generally, operably linked DNA sequences may be in close proximity or contiguous and, where necessary to join two protein coding regions, in the same reading frame.

"Peptide nucleic acid" (PNA) refers to a DNA mimic in which nucleotide bases are attached to a pseudopeptide backbone to increase stability. PNAs, also designated antigene agents, can prevent gene expression by targeting complementary messenger RNA.

The phrases "percent identity" and "% identity", as applied to polynucleotide sequences, refer to the percentage of residue matches between at least two polynucleotide sequences aligned using a standardized algorithm. Such an algorithm may insert, in a standardized and reproducible way, gaps in the sequences being compared in order to optimize alignment between two sequences, and therefore achieve a more meaningful comparison of the two sequences.

Percent identity between polynucleotide sequences may be determined using the default parameters of the CLUSTAL V algorithm as incorporated into the MEGALIGN version 3.12e sequence alignment program. This program is part of the LASERGENE software package, a suite of molecular biological analysis programs (DNASTAR, Madison WI). CLUSTAL V is described in Higgins, D.G. and Sharp, P.M. (1989) CABIOS 5:151-153 and in Higgins, D.G. et al. (1992) CABIOS 8:189-191. For pairwise alignments of polynucleotide sequences, the default parameters are set as follows: Ktuple=2, gap penalty=5, window=4, and "diagonals saved"=4. The "weighted" residue weight table is selected as the default. Percent identity is reported by CLUSTAL V as the "percent similarity" between aligned polynucleotide sequence pairs.

Alternatively, a suite of commonly used and freely available sequence comparison algorithms can be used that are provided by the National Center for Biotechnology Information (NCBI) Basic Local Alignment Search Tool (BLAST) (Altschul, S.F. et al. (1990) J. Mol. Biol. 215:403-410), which is available from several sources, including the NCBI, Bethesda, MD, and on the Internet at http://www.ncbi.nlm.nih.gov/BLAST/. The BLAST software suite includes various sequence analysis programs including "BLASTN," that is used to determine alignment between a known polynucleotide sequence and other sequences on a variety of databases. Also available is a tool called "BLAST 2 Sequences" that is used for direct pairwise comparison of two nucleotide sequences. "BLAST 2 Sequences" can be accessed and used interactively at

http://www.ncbi.nlm.nih.gov/gorf/bl2/. The "BLAST 2 Sequences" tool can be used for both BLASTN and BLASTP (discussed below). BLAST programs are commonly used with gap and other parameters set to default settings. For example, to compare two nucleotide sequences, one may use BLASTN with the "BLAST 2 Sequences" tool Version 2.0.9 (May-07-1999) set at default parameters. Such default parameters may be, for example:

Matrix: BLOSUM62

Reward for match: 1

Penalty for mismatch: -2

Open Gap: 5 and Extension Gap: 2 penalties

Gap x drop-off: 50

Expect: 10

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Percent identity may be measured over the length of an entire defined sequence, for example, as defined by a particular SEQ ID number, or may be measured over a shorter length, for example, over the length of a fragment taken from a larger, defined sequence, for instance, a fragment of at least 20, at least 30, at least 40, at least 50, at least 70, at least 100, or at least 200 contiguous nucleotides. Such lengths are exemplary only, and it is understood that any fragment length supported by the sequences shown herein, in figures or Sequence Listings, may be used to describe a length over which percentage identity may be measured.

Nucleic acid sequences that do not show a high degree of identity may nevertheless encode similar amino acid sequences due to the degeneracy of the genetic code. It is understood that changes in nucleic acid sequence can be made using this degeneracy to produce multiple nucleic acid sequences that all encode substantially the same protein.

The phrases "percent identity" and "% identity", as applied to polypeptide sequences, refer to the percentage of residue matches between at least two polypeptide sequences aligned using a

standardized algorithm. Methods of polypeptide sequence alignment are well-known. Some alignment methods take into account conservative amino acid substitutions. Such conservative substitutions, explained in more detail above, generally preserve the hydrophobicity and acidity of the substituted residue, thus preserving the structure (and therefore function) of the folded polypeptide.

Percent identity between polypeptide sequences may be determined using the default parameters of the CLUSTAL V algorithm as incorporated into the MEGALIGN version 3.12e sequence alignment program (described and referenced above). For pairwise alignments of polypeptide sequences using CLUSTAL V, the default parameters are set as follows: Ktuple=1, gap penalty=3, window=5, and "diagonals saved"=5. The PAM250 matrix is selected as the default residue weight table. As with polynucleotide alignments, the percent identity is reported by CLUSTAL V as the "percent similarity" between aligned polypeptide sequence pairs.

Alternatively the NCBI BLAST software suite may be used. For example, for a pairwise comparison of two polypeptide sequences, one may use the "BLAST 2 Sequences" tool Version 2.0.9 (May-07-1999) with BLASTP set at default parameters. Such default parameters may be, for example:

Matrix: BLOSUM62

Open Gap: 11 and Extension Gap: 1 penalty

Gap x drop-off: 50

Expect: 10

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Filter: on

Percent identity may be measured over the length of an entire defined polypeptide sequence, for example, as defined by a particular SEQ ID number, or may be measured over a shorter length, for example, over the length of a fragment taken from a larger, defined polypeptide sequence, for instance, a fragment of at least 15, at least 20, at least 30, at least 40, at least 50, at least 70 or at least 150 contiguous residues. Such lengths are exemplary only, and it is understood that any fragment length supported by the sequences shown herein, in figures or Sequence Listings, may be used to describe a length over which percentage identity may be measured.

"Post-translational modification" of an MDDT may involve lipidation, glycosylation, phosphorylation, acetylation, racemization, proteolytic cleavage, and other modifications known in the art. These processes may occur synthetically or biochemically. Biochemical modifications will vary by cell type depending on the enzymatic milieu and the MDDT.

"Probe" refers to mddt or fragments thereof, which are used to detect identical, allelic or related nucleic acid sequences. Probes are isolated oligonucleotides or polynucleotides attached to a detectable label or reporter molecule. Typical labels include radioactive isotopes, ligands,

chemiluminescent agents, and enzymes. "Primers" are short nucleic acids, usually DNA oligonucleotides, which may be annealed to a target polynucleotide by complementary base-pairing. The primer may then be extended along the target DNA strand by a DNA polymerase enzyme. Primer pairs can be used for amplification (and identification) of a nucleic acid sequence, e.g., by the polymerase chain reaction (PCR).

Probes and primers as used in the present invention typically comprise at least 15 contiguous nucleotides of a known sequence. In order to enhance specificity, longer probes and primers may also be employed, such as probes and primers that comprise at least 20, 30, 40, 50, 60, 70, 80, 90, 100, or at least 150 consecutive nucleotides of the disclosed nucleic acid sequences. Probes and primers may be considerably longer than these examples, and it is understood that any length supported by the specification, including the figures and Sequence Listing, may be used.

Methods for preparing and using probes and primers are described in the references, for example Sambrook, J. et al., (1989, Molecular Cloning: A Laboratory Manual, 2nd ed., vol. 1-3, Cold Spring Harbor Press, Plainview NY); Ausubel, F.M. et al., (1999, Short Protocols in Molecular Biology, 4th ed. Greene Publ. John Wiley & Sons Assoc. & Wiley-Intersciences, New York NY); and Innis, M. et al., (1990; PCR Protocols, A Guide to Methods and Applications, Academic Press, San Diego CA). PCR primer pairs can be derived from a known sequence, for example, by using computer programs intended for that purpose such as Primer (Version 0.5, 1991, Whitehead Institute for Biomedical Research, Cambridge MA).

Oligonucleotides for use as primers are selected using software known in the art for such purpose. For example, OLIGO 4.06 software is useful for the selection of PCR primer pairs of up to 100 nucleotides each, and for the analysis of oligonucleotides and larger polynucleotides of up to 5,000 nucleotides from an input polynucleotide sequence of up to 32 kilobases. Similar primer selection programs have incorporated additional features for expanded capabilities. For example, the PrimOU primer selection program (available to the public from the Genome Center at University of Texas South West Medical Center, Dallas TX) is capable of choosing specific primers from megabase sequences and is thus useful for designing primers on a genome-wide scope. The Primer3 primer selection program (available to the public from the Whitehead Institute/MIT Center for Genome Research, Cambridge MA) allows the user to input a "mispriming library," in which sequences to avoid as primer binding sites are user-specified. Primer3 is useful, in particular, for the selection of oligonucleotides for microarrays. (The source code for the latter two primer selection programs may also be obtained from their respective sources and modified to meet the user's specific needs.) The PrimeGen program (available to the public from the UK Human Genome Mapping Project Resource Centre, Cambridge UK) designs primers based on multiple sequence alignments, thereby allowing selection of primers that hybridize to either the most conserved or least conserved

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regions of aligned nucleic acid sequences. Hence, this program is useful for identification of both unique and conserved oligonucleotides and polynucleotide fragments. The oligonucleotides and polynucleotide fragments identified by any of the above selection methods are useful in hybridization technologies, for example, as PCR or sequencing primers, microarray elements, or specific probes to identify fully or partially complementary polynucleotides in a sample of nucleic acids. Methods of oligonucleotide selection are not limited to those described above.

"Purified" refers to molecules, either polynucleotides or polypeptides that are isolated or separated from their natural environment and are at least 60% free, preferably at least 75% free, and most preferably at least 90% free from other compounds with which they are naturally associated.

A "recombinant nucleic acid" is a sequence that is not naturally occurring or has a sequence that is made by an artificial combination of two or more otherwise separated segments of sequence. This artificial combination is often accomplished by chemical synthesis or, more commonly, by the artificial manipulation of isolated segments of nucleic acids, e.g., by genetic engineering techniques such as those described in Sambrook, <u>supra</u>. The term recombinant includes nucleic acids that have been altered solely by addition, substitution, or deletion of a portion of the nucleic acid. Frequently, a recombinant nucleic acid may include a nucleic acid sequence operably linked to a promoter sequence. Such a recombinant nucleic acid may be part of a vector that is used, for example, to transform a cell.

Alternatively, such recombinant nucleic acids may be part of a viral vector, e.g., based on a vaccinia virus, that could be use to vaccinate a mammal wherein the recombinant nucleic acid is expressed, inducing a protective immunological response in the mammal.

"Regulatory element" refers to a nucleic acid sequence from nontranslated regions of a gene, and includes enhancers, promoters, introns, and 3' untranslated regions, which interact with host proteins to carry out or regulate transcription or translation.

"Reporter" molecules are chemical or biochemical moieties used for labeling a nucleic acid, an amino acid, or an antibody. They include radionuclides; enzymes; fluorescent, chemiluminescent, or chromogenic agents; substrates; cofactors; inhibitors; magnetic particles; and other moieties known in the art.

An "RNA equivalent," in reference to a DNA sequence, is composed of the same linear sequence of nucleotides as the reference DNA sequence with the exception that all occurrences of the nitrogenous base thymine are replaced with uracil, and the sugar backbone is composed of ribose instead of deoxyribose.

"Sample" is used in its broadest sense. Samples may contain nucleic or amino acids, antibodies, or other materials, and may be derived from any source (e.g., bodily fluids including, but not limited to, saliva, blood, and urine; chromosome(s), organelles, or membranes isolated from a

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cell; genomic DNA, RNA, or cDNA in solution or bound to a substrate; and cleared cells or tissues or blots or imprints from such cells or tissues).

"Specific binding" or "specifically binding" refers to the interaction between a protein or peptide and its agonist, antibody, antagonist, or other binding partner. The interaction is dependent upon the presence of a particular structure of the protein, e.g., the antigenic determinant or epitope, recognized by the binding molecule. For example, if an antibody is specific for epitope "A," the presence of a polypeptide containing epitope A, or the presence of free unlabeled A, in a reaction containing free labeled A and the antibody will reduce the amount of labeled A that binds to the antibody.

"Substitution" refers to the replacement of at least one nucleotide or amino acid by a different nucleotide or amino acid.

"Substrate" refers to any suitable rigid or semi-rigid support including, e.g., membranes, filters, chips, slides, wafers, fibers, magnetic or nonmagnetic beads, gels, tubing, plates, polymers, microparticles or capillaries. The substrate can have a variety of surface forms, such as wells, trenches, pins, channels and pores, to which polynucleotides or polypeptides are bound.

A "transcript image" or "expression profile" refers to the collective pattern of gene expression by a particular cell type or tissue under given conditions at a given time.

"Transformation" refers to a process by which exogenous DNA enters a recipient cell.

Transformation may occur under natural or artificial conditions using various methods well known in the art. Transformation may rely on any known method for the insertion of foreign nucleic acid sequences into a prokaryotic or eukaryotic host cell. The method is selected based on the host cell being transformed.

"Transformants" include stably transformed cells in which the inserted DNA is capable of replication either as an autonomously replicating plasmid or as part of the host chromosome, as well as cells which transiently express inserted DNA or RNA.

A "transgenic organism," as used herein, is any organism, including but not limited to animals and plants, in which one or more of the cells of the organism contains heterologous nucleic acid introduced by way of human intervention, such as by transgenic techniques well known in the art. The nucleic acid is introduced into the cell, directly or indirectly by introduction into a precursor of the cell, by way of deliberate genetic manipulation, such as by microinjection or by infection with a recombinant virus. The term genetic manipulation does not include classical cross-breeding, or in vitro fertilization, but rather is directed to the introduction of a recombinant DNA molecule. The transgenic organisms contemplated in accordance with the present invention include bacteria, cyanobacteria, fungi, and plants and animals. The isolated DNA of the present invention can be introduced into the host by methods known in the art, for example infection, transfection,

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transformation or transconjugation. Techniques for transferring the DNA of the present invention into such organisms are widely known and provided in references such as Sambrook et al. (1989), <u>supra</u>.

A "variant" of a particular nucleic acid sequence is defined as a nucleic acid sequence having at least 25% sequence identity to the particular nucleic acid sequence over a certain length of one of the nucleic acid sequences using BLASTN with the "BLAST 2 Sequences" tool Version 2.0.9 (May-07-1999) set at default parameters. Such a pair of nucleic acids may show, for example, at least 30%, at least 50%, at least 60%, at least 70%, at least 80%, at least 90%, at least 91%, at least 92%, at least 93%, at least 94%, at least 95%, at least 96%, at least 97%, at least 98%, or at least 99% or greater sequence identity over a certain defined length. The variant may result in "conservative" amino acid changes which do not affect structural and/or chemical properties. A variant may be described as, for example, an "allelic" (as defined above), "splice," "species," or "polymorphic" variant. A splice variant may have significant identity to a reference molecule, but will generally have a greater or lesser number of polynucleotides due to alternate splicing of exons during mRNA processing. The corresponding polypeptide may possess additional functional domains or lack domains that are present in the reference molecule. Species variants are polynucleotide sequences that vary from one species to another. The resulting polypeptides generally will have significant amino acid identity relative to each other. A polymorphic variant is a variation in the polynucleotide sequence of a particular gene between individuals of a given species. Polymorphic variants also may encompass "single nucleotide polymorphisms" (SNPs) in which the polynucleotide sequence varies by one base. The presence of SNPs may be indicative of, for example, a certain population, a disease state, or a propensity for a disease state.

In an alternative, variants of the polynucleotides of the present invention may be generated through recombinant methods. One possible method is a DNA shuffling technique such as MOLECULARBREEDING (Maxygen Inc., Santa Clara CA; described in U.S. Patent Number 5,837,458; Chang, C.-C. et al. (1999) Nat. Biotechnol. 17:793-797; Christians, F.C. et al. (1999) Nat. Biotechnol. 17:259-264; and Crameri, A. et al. (1996) Nat. Biotechnol. 14:315-319) to alter or improve the biological properties of MDDT, such as its biological or enzymatic activity or its ability to bind to other molecules or compounds. DNA shuffling is a process by which a library of gene variants is produced using PCR-mediated recombination of gene fragments. The library is then subjected to selection or screening procedures that identify those gene variants with the desired properties. These preferred variants may then be pooled and further subjected to recursive rounds of DNA shuffling and selection/screening. Thus, genetic diversity is created through "artificial" breeding and rapid molecular evolution. For example, fragments of a single gene containing random point mutations may be recombined, screened, and then reshuffled until the desired properties are

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optimized. Alternatively, fragments of a given gene may be recombined with fragments of homologous genes in the same gene family, either from the same or different species, thereby maximizing the genetic diversity of multiple naturally occurring genes in a directed and controllable manner.

A "variant" of a particular polypeptide sequence is defined as a polypeptide sequence having at least 40% sequence identity to the particular polypeptide sequence over a certain length of one of the polypeptide sequences using BLASTP with the "BLAST 2 Sequences" tool Version 2.0.9 (May-07-1999) set at default parameters. Such a pair of polypeptides may show, for example, at least 50%, at least 60%, at least 70%, at least 80%, at least 90%, at least 91%, at least 92%, at least 93%, at least 94%, at least 95%, at least 96%, at least 97%, at least 98%, or at least 99% or greater sequence identity over a certain defined length of one of the polypeptides.

THE INVENTION

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In a particular embodiment, cDNA sequences derived from human tissues and cell lines were aligned based on nucleotide sequence identity and assembled into "consensus" or "template" sequences which are designated by the template identification numbers (template IDs) in column 2 of Table 2. The sequence identification numbers (SEQ ID NO:s) corresponding to the template IDs are shown in column 1. The template sequences have similarity to GenBank sequences, or "hits," as designated by the GI Numbers in column 3. The statistical probability of each GenBank hit is indicated by a probability score in column 4, and the functional annotation corresponding to each GenBank hit is listed in column 5.

The invention incorporates the nucleic acid sequences of these templates as disclosed in the Sequence Listing and the use of these sequences in the diagnosis and treatment of disease states characterized by defects in disease detection and treatment molecules. The invention further utilizes these sequences in hybridization and amplification technologies, and in particular, in technologies which assess gene expression patterns correlated with specific cells or tissues and their responses in vivo or in vitro to pharmaceutical agents, toxins, and other treatments. In this manner, the sequences of the present invention are used to develop a transcript image for a particular cell or tissue.

30 Derivation of Nucleic Acid Sequences

cDNA was isolated from libraries constructed using RNA derived from normal and diseased human tissues and cell lines. The human tissues and cell lines used for cDNA library construction were selected from a broad range of sources to provide a diverse population of cDNAs representative of gene transcription throughout the human body. Descriptions of the human tissues and cell lines used for cDNA library construction are provided in the LIFESEQ database (Incyte Genomics, Inc.

(Incyte), Palo Alto CA). Human tissues were broadly selected from, for example, cardiovascular, dermatologic, endocrine, gastrointestinal, hematopoietic/immune system, musculoskeletal, neural, reproductive, and urologic sources.

Cell lines used for cDNA library construction were derived from, for example, leukemic cells, teratocarcinomas, neuroepitheliomas, cervical carcinoma, lung fibroblasts, and endothelial cells. Such cell lines include, for example, THP-1, Jurkat, HUVEC, hNT2, WI38, HeLa, and other cell lines commonly used and available from public depositories (American Type Culture Collection, Manassas VA). Prior to mRNA isolation, cell lines were untreated, treated with a pharmaceutical agent such as 5'-aza-2'-deoxycytidine, treated with an activating agent such as lipopolysaccharide in the case of leukocytic cell lines, or, in the case of endothelial cell lines, subjected to shear stress.

Sequencing of the cDNAs

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Methods for DNA sequencing are well known in the art. Conventional enzymatic methods employ the Klenow fragment of DNA polymerase I, SEQUENASE DNA polymerase (U.S. Biochemical Corporation, Cleveland OH), Taq polymerase (Applied Biosystems, Foster City CA), thermostable T7 polymerase (Amersham Pharmacia Biotech, Inc. (Amersham Pharmacia Biotech), Piscataway NJ), or combinations of polymerases and proofreading exonucleases such as those found in the ELONGASE amplification system (Life Technologies Inc. (Life Technologies), Gaithersburg MD), to extend the nucleic acid sequence from an oligonucleotide primer annealed to the DNA template of interest. Methods have been developed for the use of both single-stranded and doublestranded templates. Chain termination reaction products may be electrophoresed on ureapolyacrylamide gels and detected either by autoradiography (for radioisotope-labeled nucleotides) or by fluorescence (for fluorophore-labeled nucleotides). Automated methods for mechanized reaction preparation, sequencing, and analysis using fluorescence detection methods have been developed. Machines used to prepare cDNAs for sequencing can include the MICROLAB 2200 liquid transfer system (Hamilton Company (Hamilton), Reno NV), Peltier thermal cycler (PTC200; MJ Research, Inc. (MJ Research), Watertown MA), and ABI CATALYST 800 thermal cycler (Applied Biosystems). Sequencing can be carried out using, for example, the ABI 373 or 377 (Applied Biosystems) or MEGABACE 1000 (Molecular Dynamics, Inc. (Molecular Dynamics), Sunnyvale CA) DNA sequencing systems, or other automated and manual sequencing systems well known in the art.

The nucleotide sequences of the Sequence Listing have been prepared by current, state-ofthe-art, automated methods and, as such, may contain occasional sequencing errors or unidentified nucleotides. Such unidentified nucleotides are designated by an N. These infrequent unidentified bases do not represent a hindrance to practicing the invention for those skilled in the art. Several

methods employing standard recombinant techniques may be used to correct errors and complete the missing sequence information. (See, e.g., those described in Ausubel, F.M. et al. (1997) Short Protocols in Molecular Biology, John Wiley & Sons, New York NY; and Sambrook, J. et al. (1989) Molecular Cloning, A Laboratory Manual, Cold Spring Harbor Press, Plainview NY.)

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Assembly of cDNA Sequences

Human polynucleotide sequences may be assembled using programs or algorithms well known in the art. Sequences to be assembled are related, wholly or in part, and may be derived from a single or many different transcripts. Assembly of the sequences can be performed using such programs as PHRAP (Phils Revised Assembly Program) and the GELVIEW fragment assembly system (GCG), or other methods known in the art.

Alternatively, cDNA sequences are used as "component" sequences that are assembled into "template" or "consensus" sequences as follows. Sequence chromatograms are processed, verified, and quality scores are obtained using PHRED. Raw sequences are edited using an editing pathway known as Block 1 (See, e.g., the LIFESEQ Assembled User Guide, Incyte Genomics, Palo Alto, CA). A series of BLAST comparisons is performed and low-information segments and repetitive elements (e.g., dinucleotide repeats, Alu repeats, etc.) are replaced by "n's", or masked, to prevent spurious matches. Mitochondrial and ribosomal RNA sequences are also removed. The processed sequences are then loaded into a relational database management system (RDMS) which assigns edited sequences to existing templates, if available. When additional sequences are added into the RDMS, a process is initiated which modifies existing templates or creates new templates from works in progress (i.e., nonfinal assembled sequences) containing queued sequences or the sequences themselves. After the new sequences have been assigned to templates, the templates can be merged into bins. If multiple templates exist in one bin, the bin can be split and the templates reannotated.

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Once gene bins have been generated based upon sequence alignments, bins are "clone joined" based upon clone information. Clone joining occurs when the 5' sequence of one clone is present in one bin and the 3' sequence from the same clone is present in a different bin, indicating that the two bins should be merged into a single bin. Only bins which share at least two different clones are merged.

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A resultant template sequence may contain either a partial or a full length open reading frame, or all or part of a genetic regulatory element. This variation is due in part to the fact that the full length cDNAs of many genes are several hundred, and sometimes several thousand, bases in length. With current technology, cDNAs comprising the coding regions of large genes cannot be cloned because of vector limitations, incomplete reverse transcription of the mRNA, or incomplete "second strand" synthesis. Template sequences may be extended to include additional contiguous

sequences derived from the parent RNA transcript using a variety of methods known to those of skill in the art. Extension may thus be used to achieve the full length coding sequence of a gene.

Analysis of the cDNA Sequences

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The cDNA sequences are analyzed using a variety of programs and algorithms which are well known in the art. (See, e.g., Ausubel, 1997, <u>supra</u>, Chapter 7.7; Meyers, R.A. (Ed.) (1995) <u>Molecular Biology and Biotechnology</u>, Wiley VCH, New York NY, pp. 856-853; and Table 8.) These analyses comprise both reading frame determinations, e.g., based on triplet codon periodicity for particular organisms (Fickett, J.W. (1982) Nucleic Acids Res. 10:5303-5318); analyses of potential start and stop codons; and homology searches.

Computer programs known to those of skill in the art for performing computer-assisted searches for amino acid and nucleic acid sequence similarity, include, for example, Basic Local Alignment Search Tool (BLAST; Altschul, S.F. (1993) J. Mol. Evol. 36:290-300; Altschul, S.F. et al. (1990) J. Mol. Biol. 215:403-410). BLAST is especially useful in determining exact matches and comparing two sequence fragments of arbitrary but equal lengths, whose alignment is locally maximal and for which the alignment score meets or exceeds a threshold or cutoff score set by the user (Karlin, S. et al. (1988) Proc. Natl. Acad. Sci. USA 85:841-845). Using an appropriate search tool (e.g., BLAST or HMM), GenBank, SwissProt, BLOCKS, PFAM and other databases may be searched for sequences containing regions of homology to a query mddt or MDDT of the present invention.

Other approaches to the identification, assembly, storage, and display of nucleotide and polypeptide sequences are provided in "Relational Database for Storing Biomolecule Information," U.S.S.N. 08/947,845, filed October 9, 1997; "Project-Based Full-Length Biomolecular Sequence Database," U.S. Patent Number 5,953,727; and "Relational Database and System for Storing Information Relating to Biomolecular Sequences," U.S.S.N. 09/034,807, filed March 4, 1998, all of which are incorporated by reference herein in their entirety.

Protein hierarchies can be assigned to the putative encoded polypeptide based on, e.g., motif, BLAST, or biological analysis. Methods for assigning these hierarchies are described, for example, in "Database System Employing Protein Function Hierarchies for Viewing Biomolecular Sequence Data," U.S. Patent Number 6,023,659, incorporated herein by reference.

Human Disease Detection and Treatment Molecule Sequences

The mddt of the present invention may be used for a variety of diagnostic and therapeutic purposes. For example, an mddt may be used to diagnose a particular condition, disease, or disorder associated with disease detection and treatment molecules. Such conditions, diseases, and disorders

include, but are not limited to, a cell proliferative disorder, such as actinic keratosis, arteriosclerosis, atherosclerosis, bursitis, cirrhosis, hepatitis, mixed connective tissue disease (MCTD), myelofibrosis, paroxysmal nocturnal hemoglobinuria, polycythemia vera, psoriasis, primary thrombocythemia, and cancers including adenocarcinoma, leukemia, lymphoma, melanoma, myeloma, sarcoma, teratocarcinoma, and, in particular, a cancer of the adrenal gland, bladder, bone, bone marrow, brain, breast, cervix, gall bladder, ganglia, gastrointestinal tract, heart, kidney, liver, lung, muscle, ovary, pancreas, parathyroid, penis, prostate, salivary glands, skin, spleen, testis, thymus, thyroid, and uterus; and an autoimmune/inflammatory disorder, such as actinic keratosis, acquired immunodeficiency syndrome (AIDS), Addison's disease, adult respiratory distress syndrome, allergies, ankylosing spondylitis, amyloidosis, anemia, arteriosclerosis, asthma, atherosclerosis, 10 autoimmune hemolytic anemia, autoimmune thyroiditis, bronchitis, bursitis, cholecystitis, cirrhosis, contact dermatitis, Crohn's disease, atopic dermatitis, dermatomyositis, diabetes mellitus, emphysema, erythroblastosis fetalis, erythema nodosum, atrophic gastritis, glomerulonephritis, Goodpasture's syndrome, gout, Graves' disease, Hashimoto's thyroiditis, paroxysmal nocturnal hemoglobinuria, hepatitis, hypereosinophilia, irritable bowel syndrome, episodic lymphopenia with 15 lymphocytotoxins, mixed connective tissue disease (MCTD), multiple sclerosis, myasthenia gravis, myocardial or pericardial inflammation, myelofibrosis, osteoarthritis, osteoporosis, pancreatitis, polycythemia vera, polymyositis, psoriasis, Reiter's syndrome, rheumatoid arthritis, scleroderma, Sjögren's syndrome, systemic anaphylaxis, systemic lupus erythematosus, systemic sclerosis, primary thrombocythemia, thrombocytopenic purpura, ulcerative colitis, uveitis, Werner syndrome, 20 complications of cancer, hemodialysis, and extracorporeal circulation, trauma, and hematopoietic cancer including lymphoma, leukemia, and myeloma. The mddt can be used to detect the presence of, or to quantify the amount of, an mddt-related polynucleotide in a sample. This information is then compared to information obtained from appropriate reference samples, and a diagnosis is established. Alternatively, a polynucleotide complementary to a given mddt can inhibit or inactivate a 25 therapeutically relevant gene related to the mddt.

Analysis of mddt Expression Patterns

The expression of mddt may be routinely assessed by hybridization-based methods to determine, for example, the tissue-specificity, disease-specificity, or developmental stage-specificity of mddt expression. For example, the level of expression of mddt may be compared among different cell types or tissues, among diseased and normal cell types or tissues, among cell types or tissues at different developmental stages, or among cell types or tissues undergoing various treatments. This type of analysis is useful, for example, to assess the relative levels of mddt expression in fully or partially differentiated cells or tissues, to determine if changes in mddt expression levels are

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correlated with the development or progression of specific disease states, and to assess the response of a cell or tissue to a specific therapy, for example, in pharmacological or toxicological studies. Methods for the analysis of mddt expression are based on hybridization and amplification technologies and include membrane-based procedures such as northern blot analysis, high-throughput procedures that utilize, for example, microarrays, and PCR-based procedures.

Hybridization and Genetic Analysis

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The mddt, their fragments, or complementary sequences, may be used to identify the presence of and/or to determine the degree of similarity between two (or more) nucleic acid sequences. The mddt may be hybridized to naturally occurring or recombinant nucleic acid sequences under appropriately selected temperatures and salt concentrations. Hybridization with a probe based on the nucleic acid sequence of at least one of the mddt allows for the detection of nucleic acid sequences, including genomic sequences, which are identical or related to the mddt of the Sequence Listing. Probes may be selected from non-conserved or unique regions of at least one of the polynucleotides of SEQ ID NO:1-104 and tested for their ability to identify or amplify the target nucleic acid sequence using standard protocols.

Polynucleotide sequences that are capable of hybridizing, in particular, to those shown in SEQ ID NO:1-104 and fragments thereof, can be identified using various conditions of stringency. (See, e.g., Wahl, G.M. and S.L. Berger (1987) Methods Enzymol. 152:399-407; Kimmel, A.R. (1987) Methods Enzymol. 152:507-511.) Hybridization conditions are discussed in "Definitions."

A probe for use in Southern or northern hybridization may be derived from a fragment of an mddt sequence, or its complement, that is up to several hundred nucleotides in length and is either single-stranded or double-stranded. Such probes may be hybridized in solution to biological materials such as plasmids, bacterial, yeast, or human artificial chromosomes, cleared or sectioned tissues, or to artificial substrates containing mddt. Microarrays are particularly suitable for identifying the presence of and detecting the level of expression for multiple genes of interest by examining gene expression correlated with, e.g., various stages of development, treatment with a drug or compound, or disease progression. An array analogous to a dot or slot blot may be used to arrange and link polynucleotides to the surface of a substrate using one or more of the following: mechanical (vacuum), chemical, thermal, or UV bonding procedures. Such an array may contain any number of mddt and may be produced by hand or by using available devices, materials, and machines.

Microarrays may be prepared, used, and analyzed using methods known in the art. (See, e.g., Brennan, T.M. et al. (1995) U.S. Patent No. 5,474,796; Schena, M. et al. (1996) Proc. Natl. Acad. Sci. USA 93:10614-10619; Baldeschweiler et al. (1995) PCT application WO95/251116; Shalon, D. et al. (1995) PCT application WO95/35505; Heller, R.A. et al. (1997) Proc. Natl. Acad. Sci. USA 94:2150-

2155; and Heller, M.J. et al. (1997) U.S. Patent No. 5,605,662.)

Probes may be labeled by either PCR or enzymatic techniques using a variety of commercially available reporter molecules. For example, commercial kits are available for radioactive and chemiluminescent labeling (Amersham Pharmacia Biotech) and for alkaline phosphatase labeling (Life Technologies). Alternatively, mddt may be cloned into commercially available vectors for the production of RNA probes. Such probes may be transcribed in the presence of at least one labeled nucleotide (e.g., ³²P-ATP, Amersham Pharmacia Biotech).

Additionally the polynucleotides of SEQ ID NO:1-104 or suitable fragments thereof can be used to isolate full length cDNA sequences utilizing hybridization and/or amplification procedures well known in the art, e.g., cDNA library screening, PCR amplification, etc. The molecular cloning of such full length cDNA sequences may employ the method of cDNA library screening with probes using the hybridization, stringency, washing, and probing strategies described above and in Ausubel, supra, Chapters 3, 5, and 6. These procedures may also be employed with genomic libraries to isolate genomic sequences of mddt in order to analyze, e.g., regulatory elements.

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Genetic Mapping

Gene identification and mapping are important in the investigation and treatment of almost all conditions, diseases, and disorders. Cancer, cardiovascular disease, Alzheimer's disease, arthritis, diabetes, and mental illnesses are of particular interest. Each of these conditions is more complex than the single gene defects of sickle cell anemia or cystic fibrosis, with select groups of genes being predictive of predisposition for a particular condition, disease, or disorder. For example, cardiovascular disease may result from malfunctioning receptor molecules that fail to clear cholesterol from the bloodstream, and diabetes may result when a particular individual's immune system is activated by an infection and attacks the insulin-producing cells of the pancreas. In some studies, Alzheimer's disease has been linked to a gene on chromosome 21; other studies predict a different gene and location. Mapping of disease genes is a complex and reiterative process and generally proceeds from genetic linkage analysis to physical mapping.

As a condition is noted among members of a family, a genetic linkage map traces parts of chromosomes that are inherited in the same pattern as the condition. Statistics link the inheritance of particular conditions to particular regions of chromosomes, as defined by RFLP or other markers. (See, for example, Lander, E. S. and Botstein, D. (1986) Proc. Natl. Acad. Sci. USA 83:7353-7357.) Occasionally, genetic markers and their locations are known from previous studies. More often, however, the markers are simply stretches of DNA that differ among individuals. Examples of genetic linkage maps can be found in various scientific journals or at the Online Mendelian Inheritance in Man (OMIM) World Wide Web site.

In another embodiment of the invention, mddt sequences may be used to generate hybridization probes useful in chromosomal mapping of naturally occurring genomic sequences. Either coding or noncoding sequences of mddt may be used, and in some instances, noncoding sequences may be preferable over coding sequences. For example, conservation of an mddt coding sequence among members of a multi-gene family may potentially cause undesired cross hybridization during chromosomal mapping. The sequences may be mapped to a particular chromosome, to a specific region of a chromosome, or to artificial chromosome constructions, e.g., human artificial chromosomes (HACs), yeast artificial chromosomes (YACs), bacterial artificial chromosomes (BACs), bacterial P1 constructions, or single chromosome cDNA libraries. (See, e.g., Harrington, J.J. et al. (1997) Nat. Genet. 15:345-355; Price, C.M. (1993) Blood Rev. 7:127-134; and Trask, B.J. (1991) Trends Genet. 7:149-154.)

Fluorescent in situ hybridization (FISH) may be correlated with other physical chromosome mapping techniques and genetic map data. (See, e.g., Meyers, supra, pp. 965-968.) Correlation between the location of mddt on a physical chromosomal map and a specific disorder, or a predisposition to a specific disorder, may help define the region of DNA associated with that disorder. The mddt sequences may also be used to detect polymorphisms that are genetically linked to the inheritance of a particular condition, disease, or disorder.

In situ hybridization of chromosomal preparations and genetic mapping techniques, such as linkage analysis using established chromosomal markers, may be used for extending existing genetic maps. Often the placement of a gene on the chromosome of another mammalian species, such as mouse, may reveal associated markers even if the number or arm of the corresponding human chromosome is not known. These new marker sequences can be mapped to human chromosomes and may provide valuable information to investigators searching for disease genes using positional cloning or other gene discovery techniques. Once a disease or syndrome has been crudely correlated by genetic linkage with a particular genomic region, e.g., ataxia-telangiectasia to 11q22-23, any sequences mapping to that area may represent associated or regulatory genes for further investigation. (See, e.g., Gatti, R.A. et al. (1988) Nature 336:577-580.) The nucleotide sequences of the subject invention may also be used to detect differences in chromosomal architecture due to translocation, inversion, etc., among normal, carrier, or affected individuals.

Once a disease-associated gene is mapped to a chromosomal region, the gene must be cloned in order to identify mutations or other alterations (e.g., translocations or inversions) that may be correlated with disease. This process requires a physical map of the chromosomal region containing the disease-gene of interest along with associated markers. A physical map is necessary for determining the nucleotide sequence of and order of marker genes on a particular chromosomal region. Physical mapping techniques are well known in the art and require the generation of

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overlapping sets of cloned DNA fragments from a particular organelle, chromosome, or genome. These clones are analyzed to reconstruct and catalog their order. Once the position of a marker is determined, the DNA from that region is obtained by consulting the catalog and selecting clones from that region. The gene of interest is located through positional cloning techniques using hybridization or similar methods.

Diagnostic Uses

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The mddt of the present invention may be used to design probes useful in diagnostic assays. Such assays, well known to those skilled in the art, may be used to detect or confirm conditions, disorders, or diseases associated with abnormal levels of mddt expression. Labeled probes developed from mddt sequences are added to a sample under hybridizing conditions of desired stringency. In some instances, mddt, or fragments or oligonucleotides derived from mddt, may be used as primers in amplification steps prior to hybridization. The amount of hybridization complex formed is quantified and compared with standards for that cell or tissue. If mddt expression varies significantly from the standard, the assay indicates the presence of the condition, disorder, or disease. Qualitative or quantitative diagnostic methods may include northern, dot blot, or other membrane or dip-stick based technologies or multiple-sample format technologies such as PCR, enzyme-linked immunosorbent assay (ELISA)-like, pin, or chip-based assays.

The probes described above may also be used to monitor the progress of conditions, disorders, or diseases associated with abnormal levels of mddt expression, or to evaluate the efficacy of a particular therapeutic treatment. The candidate probe may be identified from the mddt that are specific to a given human tissue and have not been observed in GenBank or other genome databases. Such a probe may be used in animal studies, preclinical tests, clinical trials, or in monitoring the treatment of an individual patient. In a typical process, standard expression is established by methods well known in the art for use as a basis of comparison, samples from patients affected by the disorder or disease are combined with the probe to evaluate any deviation from the standard profile, and a therapeutic agent is administered and effects are monitored to generate a treatment profile. Efficacy is evaluated by determining whether the expression progresses toward or returns to the standard normal pattern. Treatment profiles may be generated over a period of several days or several months. Statistical methods well known to those skilled in the art may be use to determine the significance of such therapeutic agents.

The polynucleotides are also useful for identifying individuals from minute biological samples, for example, by matching the RFLP pattern of a sample's DNA to that of an individual's DNA. The polynucleotides of the present invention can also be used to determine the actual base-by-base DNA sequence of selected portions of an individual's genome. These sequences can be

used to prepare PCR primers for amplifying and isolating such selected DNA, which can then be sequenced. Using this technique, an individual can be identified through a unique set of DNA sequences. Once a unique ID database is established for an individual, positive identification of that individual can be made from extremely small tissue samples.

In a particular aspect, oligonucleotide primers derived from the mddt of the invention may be used to detect single nucleotide polymorphisms (SNPs). SNPs are substitutions, insertions and deletions that are a frequent cause of inherited or acquired genetic disease in humans. Methods of SNP detection include, but are not limited to, single-stranded conformation polymorphism (SSCP) and fluorescent SSCP (fSSCP) methods. In SSCP, oligonucleotide primers derived from mddt are used to amplify DNA using the polymerase chain reaction (PCR). The DNA may be derived, for example, from diseased or normal tissue, biopsy samples, bodily fluids, and the like. SNPs in the DNA cause differences in the secondary and tertiary structures of PCR products in single-stranded form, and these differences are detectable using gel electrophoresis in non-denaturing gels. In fSCCP, the oligonucleotide primers are fluorescently labeled, which allows detection of the amplimers in high-throughput equipment such as DNA sequencing machines. Additionally, sequence database analysis methods, termed in silico SNP (isSNP), are capable of identifying polymorphisms by comparing the sequences of individual overlapping DNA fragments which assemble into a common consensus sequence. These computer-based methods filter out sequence variations due to laboratory preparation of DNA and sequencing errors using statistical models and automated analyses of DNA sequence chromatograms. In the alternative, SNPs may be detected and characterized by mass spectrometry using, for example, the high throughput MASSARRAY system (Sequenom, Inc., San Diego CA).

DNA-based identification techniques are critical in forensic technology. DNA sequences taken from very small biological samples such as tissues, e.g., hair or skin, or body fluids, e.g., blood, saliva, semen, etc., can be amplified using, e.g., PCR, to identify individuals. (See, e.g., Erlich, H. (1992) PCR Technology, Freeman and Co., New York, NY). Similarly, polynucleotides of the present invention can be used as polymorphic markers.

There is also a need for reagents capable of identifying the source of a particular tissue. Appropriate reagents can comprise, for example, DNA probes or primers prepared from the sequences of the present invention that are specific for particular tissues. Panels of such reagents can identify tissue by species and/or by organ type. In a similar fashion, these reagents can be used to screen tissue cultures for contamination.

The polynucleotides of the present invention can also be used as molecular weight markers on nucleic acid gels or Southern blots, as diagnostic probes for the presence of a specific mRNA in a particular cell type, in the creation of subtracted cDNA libraries which aid in the discovery of novel

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polynucleotides, in selection and synthesis of oligomers for attachment to an array or other support, and as an antigen to elicit an immune response.

Disease Model Systems Using mddt

The mddt of the invention or their mammalian homologs may be "knocked out" in an animal model system using homologous recombination in embryonic stem (ES) cells. Such techniques are well known in the art and are useful for the generation of animal models of human disease. (See, e.g., U.S. Patent Number 5,175,383 and U.S. Patent Number 5,767,337.) For example, mouse ES cells, such as the mouse 129/SvJ cell line, are derived from the early mouse embryo and grown in culture. The ES cells are transformed with a vector containing the gene of interest disrupted by a marker gene, e.g., the neomycin phosphotransferase gene (neo; Capecchi, M.R. (1989) Science 244:1288-1292). The vector integrates into the corresponding region of the host genome by homologous recombination. Alternatively, homologous recombination takes place using the Cre-loxP system to knockout a gene of interest in a tissue- or developmental stage-specific manner (Marth, J.D. (1996) Clin. Invest. 97:1999-2002; Wagner, K.U. et al. (1997) Nucleic Acids Res. 25:4323-4330). Transformed ES cells are identified and microinjected into mouse cell blastocysts such as those from the C57BL/6 mouse strain. The blastocysts are surgically transferred to pseudopregnant dams, and the resulting chimeric progeny are genotyped and bred to produce heterozygous or homozygous

The mddt of the invention may also be manipulated <u>in vitro</u> in ES cells derived from human blastocysts. Human ES cells have the potential to differentiate into at least eight separate cell lineages including endoderm, mesoderm, and ectodermal cell types. These cell lineages differentiate into, for example, neural cells, hematopoietic lineages, and cardiomyocytes (Thomson, J.A. et al. (1998) Science 282:1145-1147).

strains. Transgenic animals thus generated may be tested with potential therapeutic or toxic agents.

The mddt of the invention can also be used to create "knockin" humanized animals (pigs) or transgenic animals (mice or rats) to model human disease. With knockin technology, a region of mddt is injected into animal ES cells, and the injected sequence integrates into the animal cell genome. Transformed cells are injected into blastulae, and the blastulae are implanted as described above. Transgenic progeny or inbred lines are studied and treated with potential pharmaceutical agents to obtain information on treatment of a human disease. Alternatively, a mammal inbred to overexpress mddt, resulting, e.g., in the secretion of MDDT in its milk, may also serve as a convenient source of that protein (Janne, J. et al. (1998) Biotechnol. Annu. Rev. 4:55-74).

Screening Assays

MDDT encoded by polynucleotides of the present invention may be used to screen for molecules that bind to or are bound by the encoded polypeptides. The binding of the polypeptide and

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the molecule may activate (agonist), increase, inhibit (antagonist), or decrease activity of the polypeptide or the bound molecule. Examples of such molecules include antibodies, oligonucleotides, proteins (e.g., receptors), or small molecules.

Preferably, the molecule is closely related to the natural ligand of the polypeptide, e.g., a ligand or fragment thereof, a natural substrate, or a structural or functional mimetic. (See, Coligan et al., (1991) <u>Current Protocols in Immunology</u> 1(2): Chapter 5.) Similarly, the molecule can be closely related to the natural receptor to which the polypeptide binds, or to at least a fragment of the receptor, e.g., the active site. In either case, the molecule can be rationally designed using known techniques. Preferably, the screening for these molecules involves producing appropriate cells which express the polypeptide, either as a secreted protein or on the cell membrane. Preferred cells include cells from mammals, yeast, <u>Drosophila</u>, or <u>E. coli</u>. Cells expressing the polypeptide or cell membrane fractions which contain the expressed polypeptide are then contacted with a test compound and binding, stimulation, or inhibition of activity of either the polypeptide or the molecule is analyzed.

An assay may simply test binding of a candidate compound to the polypeptide, wherein binding is detected by a fluorophore, radioisotope, enzyme conjugate, or other detectable label. Alternatively, the assay may assess binding in the presence of a labeled competitor.

Additionally, the assay can be carried out using cell-free preparations, polypeptide/molecule affixed to a solid support, chemical libraries, or natural product mixtures. The assay may also simply comprise the steps of mixing a candidate compound with a solution containing a polypeptide, measuring polypeptide/molecule activity or binding, and comparing the polypeptide/molecule activity or binding to a standard.

Preferably, an ELISA assay using, e.g., a monoclonal or polyclonal antibody, can measure polypeptide level in a sample. The antibody can measure polypeptide level by either binding, directly or indirectly, to the polypeptide or by competing with the polypeptide for a substrate.

All of the above assays can be used in a diagnostic or prognostic context. The molecules discovered using these assays can be used to treat disease or to bring about a particular result in a patient (e.g., blood vessel growth) by activating or inhibiting the polypeptide/molecule. Moreover, the assays can discover agents which may inhibit or enhance the production of the polypeptide from suitably manipulated cells or tissues.

Transcript Imaging and Toxicological Testing

Another embodiment relates to the use of mddt to develop a transcript image of a tissue or cell type. A transcript image represents the global pattern of gene expression by a particular tissue or cell type. Global gene expression patterns are analyzed by quantifying the number of expressed genes and their relative abundance under given conditions and at a given time. (See Seilhamer et al.,

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"Comparative Gene Transcript Analysis," U.S. Patent Number 5,840,484, expressly incorporated by reference herein.) Thus a transcript image may be generated by hybridizing the polynucleotides of the present invention or their complements to the totality of transcripts or reverse transcripts of a particular tissue or cell type. In one embodiment, the hybridization takes place in high-throughput format, wherein the polynucleotides of the present invention or their complements comprise a subset of a plurality of elements on a microarray. The resultant transcript image would provide a profile of gene activity pertaining to disease detection and treatment molecules.

Transcript images which profile mddt expression may be generated using transcripts isolated from tissues, cell lines, biopsies, or other biological samples. The transcript image may thus reflect mddt expression in vivo, as in the case of a tissue or biopsy sample, or in vitro, as in the case of a cell line.

Transcript images which profile mddt expression may also be used in conjunction with in vitro model systems and preclinical evaluation of pharmaceuticals, as well as toxicological testing of industrial and naturally-occurring environmental compounds. All compounds induce characteristic gene expression patterns, frequently termed molecular fingerprints or toxicant signatures, which are indicative of mechanisms of action and toxicity (Nuwaysir, E. F. et al. (1999) Mol. Carcinog. 24:153-159; Steiner, S. and Anderson, N. L. (2000) Toxicol. Lett. 112-113:467-71, expressly incorporated by reference herein). If a test compound has a signature similar to that of a compound with known toxicity, it is likely to share those toxic properties. These fingerprints or signatures are most useful and refined when they contain expression information from a large number of genes and gene families. Ideally, a genome-wide measurement of expression provides the highest quality signature. Even genes whose expression is not altered by any tested compounds are important as well, as the levels of expression of these genes are used to normalize the rest of the expression data. The normalization procedure is useful for comparison of expression data after treatment with different compounds. While the assignment of gene function to elements of a toxicant signature aids in interpretation of toxicity mechanisms, knowledge of gene function is not necessary for the statistical matching of signatures which leads to prediction of toxicity. (See, for example, Press Release 00-02 from the National Institute of Environmental Health Sciences, released February 29, 2000, available at http://www.niehs.nih.gov/oc/news/toxchip.htm.) Therefore, it is important and desirable in toxicological screening using toxicant signatures to include all expressed gene sequences.

In one embodiment, the toxicity of a test compound is assessed by treating a biological sample containing nucleic acids with the test compound. Nucleic acids that are expressed in the treated biological sample are hybridized with one or more probes specific to the polynucleotides of the present invention, so that transcript levels corresponding to the polynucleotides of the present invention may be quantified. The transcript levels in the treated biological sample are compared with

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levels in an untreated biological sample. Differences in the transcript levels between the two samples are indicative of a toxic response caused by the test compound in the treated sample.

Another particular embodiment relates to the use of MDDT encoded by polynucleotides of the present invention to analyze the proteome of a tissue or cell type. The term proteome refers to the global pattern of protein expression in a particular tissue or cell type. Each protein component of a proteome can be subjected individually to further analysis. Proteome expression patterns, or profiles, are analyzed by quantifying the number of expressed proteins and their relative abundance under given conditions and at a given time. A profile of a cell's proteome may thus be generated by separating and analyzing the polypeptides of a particular tissue or cell type. In one embodiment, the separation is achieved using two-dimensional gel electrophoresis, in which proteins from a sample are separated by isoelectric focusing in the first dimension, and then according to molecular weight by sodium dodecyl sulfate slab gel electrophoresis in the second dimension (Steiner and Anderson, supra). The proteins are visualized in the gel as discrete and uniquely positioned spots, typically by staining the gel with an agent such as Coomassie Blue or silver or fluorescent stains. The optical density of each protein spot is generally proportional to the level of the protein in the sample. The optical densities of equivalently positioned protein spots from different samples, for example, from biological samples either treated or untreated with a test compound or therapeutic agent, are compared to identify any changes in protein spot density related to the treatment. The proteins in the spots are partially sequenced using, for example, standard methods employing chemical or enzymatic cleavage followed by mass spectrometry. The identity of the protein in a spot may be determined by comparing its partial sequence, preferably of at least 5 contiguous amino acid residues, to the polypeptide sequences of the present invention. In some cases, further sequence data may be obtained for definitive protein identification.

A proteomic profile may also be generated using antibodies specific for MDDT to quantify the levels of MDDT expression. In one embodiment, the antibodies are used as elements on a microarray, and protein expression levels are quantified by exposing the microarray to the sample and detecting the levels of protein bound to each array element (Lucking, A. et al. (1999) Anal. Biochem. 270:103-11; Mendoze, L. G. et al. (1999) Biotechniques 27:778-88). Detection may be performed by a variety of methods known in the art, for example, by reacting the proteins in the sample with a thiolor amino-reactive fluorescent compound and detecting the amount of fluorescence bound at each array element.

Toxicant signatures at the proteome level are also useful for toxicological screening, and should be analyzed in parallel with toxicant signatures at the transcript level. There is a poor correlation between transcript and protein abundances for some proteins in some tissues (Anderson, N. L. and Seilhamer, J. (1997) Electrophoresis 18:533-537), so proteome toxicant signatures may be

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useful in the analysis of compounds which do not significantly affect the transcript image, but which alter the proteomic profile. In addition, the analysis of transcripts in body fluids is difficult, due to rapid degradation of mRNA, so proteomic profiling may be more reliable and informative in such cases.

In another embodiment, the toxicity of a test compound is assessed by treating a biological sample containing proteins with the test compound. Proteins that are expressed in the treated biological sample are separated so that the amount of each protein can be quantified. The amount of each protein is compared to the amount of the corresponding protein in an untreated biological sample. A difference in the amount of protein between the two samples is indicative of a toxic response to the test compound in the treated sample. Individual proteins are identified by sequencing the amino acid residues of the individual proteins and comparing these partial sequences to the MDDT encoded by polynucleotides of the present invention.

In another embodiment, the toxicity of a test compound is assessed by treating a biological sample containing proteins with the test compound. Proteins from the biological sample are incubated with antibodies specific to the MDDT encoded by polynucleotides of the present invention. The amount of protein recognized by the antibodies is quantified. The amount of protein in the treated biological sample is compared with the amount in an untreated biological sample. A difference in the amount of protein between the two samples is indicative of a toxic response to the test compound in the treated sample.

Transcript images may be used to profile mddt expression in distinct tissue types. This process can be used to determine disease detection and treatment molecule activity in a particular tissue type relative to this activity in a different tissue type. Transcript images may be used to generate a profile of mddt expression characteristic of diseased tissue. Transcript images of tissues before and after treatment may be used for diagnostic purposes, to monitor the progression of disease, and to monitor the efficacy of drug treatments for diseases which affect the activity of disease detection and treatment molecules.

Transcript images of cell lines can be used to assess disease detection and treatment molecule activity and/or to identify cell lines that lack or misregulate this activity. Such cell lines may then be treated with pharmaceutical agents, and a transcript image following treatment may indicate the efficacy of these agents in restoring desired levels of this activity. A similar approach may be used to assess the toxicity of pharmaceutical agents as reflected by undesirable changes in disease detection and treatment molecule activity. Candidate pharmaceutical agents may be evaluated by comparing their associated transcript images with those of pharmaceutical agents of known effectiveness.

Antisense Molecules

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The polynucleotides of the present invention are useful in antisense technology. Antisense technology or therapy relies on the modulation of expression of a target protein through the specific binding of an antisense sequence to a target sequence encoding the target protein or directing its expression. (See, e.g., Agrawal, S., ed. (1996) Antisense Therapeutics, Humana Press Inc., Totawa NJ; Alama, A. et al. (1997) Pharmacol. Res. 36(3):171-178; Crooke, S.T. (1997) Adv. Pharmacol. 40:1-49; Sharma, H.W. and R. Narayanan (1995) Bioessays 17(12):1055-1063; and Lavrosky, Y. et al. (1997) Biochem. Mol. Med. 62(1):11-22.) An antisense sequence is a polynucleotide sequence capable of specifically hybridizing to at least a portion of the target sequence. Antisense sequences bind to cellular mRNA and/or genomic DNA, affecting translation and/or transcription. Antisense sequences can be DNA, RNA, or nucleic acid mimics and analogs. (See, e.g., Rossi, J.J. et al. (1991) Antisense Res. Dev. 1(3):285-288; Lee, R. et al. (1998) Biochemistry 37(3):900-1010; Pardridge, W.M. et al. (1995) Proc. Natl. Acad. Sci. USA 92(12):5592-5596; and Nielsen, P. E. and Haaima, G. (1997) Chem. Soc. Rev. 96:73-78.) Typically, the binding which results in modulation of expression occurs through hybridization or binding of complementary base pairs. Antisense sequences can also bind to DNA duplexes through specific interactions in the major groove of the double helix.

The polynucleotides of the present invention and fragments thereof can be used as antisense sequences to modify the expression of the polypeptide encoded by mddt. The antisense sequences can be produced ex vivo, such as by using any of the ABI nucleic acid synthesizer series (Applied Biosystems) or other automated systems known in the art. Antisense sequences can also be produced biologically, such as by transforming an appropriate host cell with an expression vector containing the sequence of interest. (See, e.g., Agrawal, supra.)

In therapeutic use, any gene delivery system suitable for introduction of the antisense sequences into appropriate target cells can be used. Antisense sequences can be delivered intracellularly in the form of an expression plasmid which, upon transcription, produces a sequence complementary to at least a portion of the cellular sequence encoding the target protein. (See, e.g., Slater, J.E., et al. (1998) J. Allergy Clin. Immunol. 102(3):469-475; and Scanlon, K.J., et al. (1995) 9(13):1288-1296.) Antisense sequences can also be introduced intracellularly through the use of viral vectors, such as retrovirus and adeno-associated virus vectors. (See, e.g., Miller, A.D. (1990) Blood 76:271; Ausubel, F.M. et al. (1995) Current Protocols in Molecular Biology, John Wiley & Sons, New York NY; Uckert, W. and W. Walther (1994) Pharmacol. Ther. 63(3):323-347.) Other gene delivery mechanisms include liposome-derived systems, artificial viral envelopes, and other systems known in the art. (See, e.g., Rossi, J.J. (1995) Br. Med. Bull. 51(1):217-225; Boado, R.J. et al. (1998) J. Pharm. Sci. 87(11):1308-1315; and Morris, M.C. et al. (1997) Nucleic Acids Res. 25(14):2730-2736.)

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Expression

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In order to express a biologically active MDDT, the nucleotide sequences encoding MDDT or fragments thereof may be inserted into an appropriate expression vector, i.e., a vector which contains the necessary elements for transcriptional and translational control of the inserted coding sequence in a suitable host. Methods which are well known to those skilled in the art may be used to construct expression vectors containing sequences encoding MDDT and appropriate transcriptional and translational control elements. These methods include in vitro recombinant DNA techniques, synthetic techniques, and in vivo genetic recombination. (See, e.g., Sambrook, supra, Chapters 4, 8, 16, and 17; and Ausubel, supra, Chapters 9, 10, 13, and 16.)

A variety of expression vector/host systems may be utilized to contain and express sequences encoding MDDT. These include, but are not limited to, microorganisms such as bacteria transformed with recombinant bacteriophage, plasmid, or cosmid DNA expression vectors; yeast transformed with yeast expression vectors; insect cell systems infected with viral expression vectors (e.g., baculovirus); plant cell systems transformed with viral expression vectors (e.g., cauliflower mosaic virus, CaMV, or tobacco mosaic virus, TMV) or with bacterial expression vectors (e.g., Ti or pBR322 plasmids); or animal (mammalian) cell systems. (See, e.g., Sambrook, supra; Ausubel, 1995, supra, Van Heeke, G. and S.M. Schuster (1989) J. Biol. Chem. 264:5503-5509; Bitter, G.A. et al. (1987) Methods Enzymol. 153:516-544; Scorer, C.A. et al. (1994) Bio/Technology 12:181-184; Engelhard, E.K. et al. (1994) Proc. Natl. Acad. Sci. USA 91:3224-3227; Sandig, V. et al. (1996) Hum. Gene Ther. 7:1937-1945; Takamatsu, N. (1987) EMBO J. 6:307-311; Coruzzi, G. et al. (1984) EMBO J. 3:1671-1680; Broglie, R. et al. (1984) Science 224:838-843; Winter, J. et al. (1991) Results Probl. Cell Differ. 17:85-105; The McGraw Hill Yearbook of Science and Technology (1992) McGraw Hill, New York NY, pp. 191-196; Logan, J. and T. Shenk (1984) Proc. Natl. Acad. Sci. USA 81:3655-3659; and Harrington, J.J. et al. (1997) Nat. Genet. 15:345-355.) Expression vectors derived from retroviruses, adenoviruses, or herpes or vaccinia viruses, or from various bacterial plasmids, may be used for delivery of nucleotide sequences to the targeted organ, tissue, or cell population. (See, e.g., Di Nicola, M. et al. (1998) Cancer Gen. Ther. 5(6):350-356; Yu, M. et al., (1993) Proc. Natl. Acad. Sci. USA 90(13):6340-6344; Buller, R.M. et al. (1985) Nature 317(6040):813-815; McGregor, D.P. et al. (1994) Mol. Immunol. 31(3):219-226; and Verma, I.M. and N. Somia (1997) Nature 389:239-242.) The invention is not limited by the host cell employed.

For long term production of recombinant proteins in mammalian systems, stable expression of MDDT in cell lines is preferred. For example, sequences encoding MDDT can be transformed into cell lines using expression vectors which may contain viral origins of replication and/or endogenous expression elements and a selectable marker gene on the same or on a separate vector. Any number of selection systems may be used to recover transformed cell lines. (See, e.g., Wigler,

M. et al. (1977) Cell 11:223-232; Lowy, I. et al. (1980) Cell 22:817-823.; Wigler, M. et al. (1980) Proc. Natl. Acad. Sci. USA 77:3567-3570; Colbere-Garapin, F. et al. (1981) J. Mol. Biol. 150:1-14; Hartman, S.C. and R.C.Mulligan (1988) Proc. Natl. Acad. Sci. USA 85:8047-8051; Rhodes, C.A. (1995) Methods Mol. Biol. 55:121-131.)

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Therapeutic Uses of mddt

The mddt of the invention may be used for somatic or germline gene therapy. Gene therapy may be performed to (i) correct a genetic deficiency (e.g., in the cases of severe combined immunodeficiency (SCID)-X1 disease characterized by X-linked inheritance (Cavazzana-Calvo, M. et al. (2000) Science 288:669-672), severe combined immunodeficiency syndrome associated with an inherited adenosine dearninase (ADA) deficiency (Blaese, R.M. et al. (1995) Science 270:475-480; Bordignon, C. et al. (1995) Science 270:470-475), cystic fibrosis (Zabner, J. et al. (1993) Cell 75:207-216; Crystal, R.G. et al. (1995) Hum. Gene Therapy 6:643-666; Crystal, R.G. et al. (1995) Hum. Gene Therapy 6:667-703), thalassemias, familial hypercholesterolemia, and hemophilia resulting from Factor VIII or Factor IX deficiencies (Crystal, R.G. (1995) Science 270:404-410; Verma, I.M. and Somia, N. (1997) Nature 389:239-242)), (ii) express a conditionally lethal gene product (e.g., in the case of cancers which result from unregulated cell proliferation), or (iii) express a protein which affords protection against intracellular parasites (e.g., against human retroviruses, such as human immunodeficiency virus (HIV) (Baltimore, D. (1988) Nature 335:395-396; Poeschla, E. et al. (1996) Proc. Natl. Acad. Sci. USA. 93:11395-11399), hepatitis B or C virus (HBV, HCV); fungal parasites, such as Candida albicans and Paracoccidioides brasiliensis; and protozoan parasites such as Plasmodium falciparum and Trypanosoma cruzi). In the case where a genetic deficiency in mddt expression or regulation causes disease, the expression of mddt from an appropriate population of transduced cells may alleviate the clinical manifestations caused by the genetic deficiency.

In a further embodiment of the invention, diseases or disorders caused by deficiencies in mddt are treated by constructing mammalian expression vectors comprising mddt and introducing these vectors by mechanical means into mddt-deficient cells. Mechanical transfer technologies for use with cells in vivo or ex vitro include (i) direct DNA microinjection into individual cells, (ii) ballistic gold particle delivery, (iii) liposome-mediated transfection, (iv) receptor-mediated gene transfer, and (v) the use of DNA transposons (Morgan, R.A. and Anderson, W.F. (1993) Annu. Rev. Biochem. 62:191-217; Ivics, Z. (1997) Cell 91:501-510; Boulay, J-L. and Récipon, H. (1998) Curr. Opin. Biotechnol. 9:445-450).

Expression vectors that may be effective for the expression of mddt include, but are not limited to, the PCDNA 3.1, EPITAG, PRCCMV2, PREP, PVAX vectors (Invitrogen, Carlsbad CA), PCMV-SCRIPT, PCMV-TAG, PEGSH/PERV (Stratagene, La Jolla CA), and PTET-OFF,

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PTET-ON, PTRE2, PTRE2-LUC, PTK-HYG (Clontech, Palo Alto CA). The mddt of the invention may be expressed using (i) a constitutively active promoter, (e.g., from cytomegalovirus (CMV), Rous sarcoma virus (RSV), SV40 virus, thymidine kinase (TK), or β-actin genes), (ii) an inducible promoter (e.g., the tetracycline-regulated promoter (Gossen, M. and Bujard, H. (1992) Proc. Natl. Acad. Sci. U.S.A. 89:5547-5551; Gossen, M. et al., (1995) Science 268:1766-1769; Rossi, F.M.V. and Blau, H.M. (1998) Curr. Opin. Biotechnol. 9:451-456), commercially available in the T-REX plasmid (Invitrogen); the ecdysone-inducible promoter (available in the plasmids PVGRXR and PIND; Invitrogen); the FK506/rapamycin inducible promoter; or the RU486/mifepristone inducible promoter (Rossi, F.M.V. and Blau, H.M. supra), or (iii) a tissue-specific promoter or the native promoter of the endogenous gene encoding MDDT from a normal individual. 10

Commercially available liposome transformation kits (e.g., the PERFECT LIPID TRANSFECTION KIT, available from Invitrogen) allow one with ordinary skill in the art to deliver polynucleotides to target cells in culture and require minimal effort to optimize experimental parameters. In the alternative, transformation is performed using the calcium phosphate method (Graham, F.L. and Eb, A.J. (1973) Virology 52:456-467), or by electroporation (Neumann, E. et al. (1982) EMBO J. 1:841-845). The introduction of DNA to primary cells requires modification of these standardized mammalian transfection protocols.

In another embodiment of the invention, diseases or disorders caused by genetic defects with respect to mddt expression are treated by constructing a retrovirus vector consisting of (i) mddt under the control of an independent promoter or the retrovirus long terminal repeat (LTR) promoter, (ii) appropriate RNA packaging signals, and (iii) a Rev-responsive element (RRE) along with additional retrovirus cis-acting RNA sequences and coding sequences required for efficient vector propagation. Retrovirus vectors (e.g., PFB and PFBNEO) are commercially available (Stratagene) and are based on published data (Riviere, I. et al. (1995) Proc. Natl. Acad. Sci. U.S.A. 92:6733-6737), incorporated by reference herein. The vector is propagated in an appropriate vector producing cell line (VPCL) that expresses an envelope gene with a tropism for receptors on the target cells or a promiscuous envelope protein such as VSVg (Armentano, D. et al. (1987) J. Virol. 61:1647-1650; Bender, M.A. et al. (1987) J. Virol. 61:1639-1646; Adam, M.A. and Miller, A.D. (1988) J. Virol. 62:3802-3806; Dull, T. et al. (1998) J. Virol. 72:8463-8471; Zufferey, R. et al. (1998) J. Virol. 72:9873-9880). U.S. Patent Number 5,910,434 to Rigg ("Method for obtaining retrovirus packaging cell lines producing high transducing efficiency retroviral supernatant") discloses a method for obtaining retrovirus packaging cell lines and is hereby incorporated by reference. Propagation of retrovirus vectors, transduction of a population of cells (e.g., CD4+ T-cells), and the return of transduced cells to a patient are procedures well known to persons skilled in the art of gene therapy and have been well documented (Ranga, U. et al. (1997) J. Virol. 71:7020-7029; Bauer, G. et al. (1997) Blood 89:2259-2267; 35

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Bonyhadi, M.L. (1997) J. Virol. 71:4707-4716; Ranga, U. et al. (1998) Proc. Natl. Acad. Sci. U.S.A. 95:1201-1206; Su, L. (1997) Blood 89:2283-2290).

In the alternative, an adenovirus-based gene therapy delivery system is used to deliver mddt to cells which have one or more genetic abnormalities with respect to the expression of mddt. The construction and packaging of adenovirus-based vectors are well known to those with ordinary skill in the art. Replication defective adenovirus vectors have proven to be versatile for importing genes encoding immunoregulatory proteins into intact islets in the pancreas (Csete, M.E. et al. (1995) Transplantation 27:263-268). Potentially useful adenoviral vectors are described in U.S. Patent Number 5,707,618 to Armentano ("Adenovirus vectors for gene therapy"), hereby incorporated by reference. For adenoviral vectors, see also Antinozzi, P.A. et al. (1999) Annu. Rev. Nutr. 19:511-544 and Verma, I.M. and Somia, N. (1997) Nature 18:389:239-242, both incorporated by reference herein.

In another alternative, a herpes-based, gene therapy delivery system is used to deliver mddt to target cells which have one or more genetic abnormalities with respect to the expression of mddt. The use of herpes simplex virus (HSV)-based vectors may be especially valuable for introducing mddt to cells of the central nervous system, for which HSV has a tropism. The construction and packaging of herpes-based vectors are well known to those with ordinary skill in the art. A replication-competent herpes simplex virus (HSV) type 1-based vector has been used to deliver a reporter gene to the eyes of primates (Liu, X. et al. (1999) Exp. Eye Res.169:385-395). The construction of a HSV-1 virus vector has also been disclosed in detail in U.S. Patent Number 5,804,413 to DeLuca ("Herpes simplex virus strains for gene transfer"), which is hereby incorporated by reference. U.S. Patent Number 5,804,413 teaches the use of recombinant HSV d92 which consists of a genome containing at least one exogenous gene to be transferred to a cell under the control of the appropriate promoter for purposes including human gene therapy. Also taught by this patent are the construction and use of recombinant HSV strains deleted for ICP4, ICP27 and ICP22. For HSV vectors, see also Goins, W. F. et al. 1999 J. Virol. 73:519-532 and Xu, H. et al., (1994) Dev. Biol. 163:152-161, hereby incorporated by reference. The manipulation of cloned herpesvirus sequences, the generation of recombinant virus following the transfection of multiple plasmids containing different segments of the large herpesvirus genomes, the growth and propagation of herpesvirus, and the infection of cells with herpesvirus are techniques well known to those of ordinary skill in the art.

In another alternative, an alphavirus (positive, single-stranded RNA virus) vector is used to deliver mddt to target cells. The biology of the prototypic alphavirus, Semliki Forest Virus (SFV), has been studied extensively and gene transfer vectors have been based on the SFV genome (Garoff, H. and Li, K-J. (1998) Curr. Opin. Biotech. 9:464-469). During alphavirus RNA replication, a subgenomic RNA is generated that normally encodes the viral capsid proteins. This subgenomic RNA replicates to higher levels than the full-length genomic RNA, resulting in the overproduction of

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capsid proteins relative to the viral proteins with enzymatic activity (e.g., protease and polymerase). Similarly, inserting mddt into the alphavirus genome in place of the capsid-coding region results in the production of a large number of mddt RNAs and the synthesis of high levels of MDDT in vector transduced cells. While alphavirus infection is typically associated with cell lysis within a few days, the ability to establish a persistent infection in hamster normal kidney cells (BHK-21) with a variant of Sindbis virus (SIN) indicates that the lytic replication of alphaviruses can be altered to suit the needs of the gene therapy application (Dryga, S.A. et al. (1997) Virology 228:74-83). The wide host range of alphaviruses will allow the introduction of mddt into a variety of cell types. The specific transduction of a subset of cells in a population may require the sorting of cells prior to transduction. The methods of manipulating infectious cDNA clones of alphaviruses, performing alphavirus cDNA and RNA transfections, and performing alphavirus infections, are well known to those with ordinary skill in the art.

Antibodies

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Anti-MDDT antibodies may be used to analyze protein expression levels. Such antibodies include, but are not limited to, polyclonal, monoclonal, chimeric, single chain, and Fab fragments. For descriptions of and protocols of antibody technologies, see, e.g., Pound J.D. (1998)

Immunochemical Protocols, Humana Press, Totowa, NJ.

The amino acid sequence encoded by the mddt of the Sequence Listing may be analyzed by appropriate software (e.g., LASERGENE NAVIGATOR software, DNASTAR) to determine regions of high immunogenicity. The optimal sequences for immunization are selected from the C-terminus, the N-terminus, and those intervening, hydrophilic regions of the polypeptide which are likely to be exposed to the external environment when the polypeptide is in its natural conformation. Analysis used to select appropriate epitopes is also described by Ausubel (1997, supra, Chapter 11.7).

Peptides used for antibody induction do not need to have biological activity; however, they must be antigenic. Peptides used to induce specific antibodies may have an amino acid sequence consisting of at least five amino acids, preferably at least 10 amino acids, and most preferably at least 15 amino acids. A peptide which mimics an antigenic fragment of the natural polypeptide may be fused with another protein such as keyhole hemolimpet cyanin (KLH; Sigma, St. Louis MO) for antibody production. A peptide encompassing an antigenic region may be expressed from an mddt, synthesized as described above, or purified from human cells.

Procedures well known in the art may be used for the production of antibodies. Various hosts including mice, goats, and rabbits, may be immunized by injection with a peptide. Depending on the host species, various adjuvants may be used to increase immunological response.

In one procedure, peptides about 15 residues in length may be synthesized using an ABI 431A peptide synthesizer (Applied Biosystems) using fmoc-chemistry and coupled to KLH (Sigma) by reaction with M-maleimidobenzoyl-N-hydroxysuccinimide ester (Ausubel, 1995, supra). Rabbits are immunized with the peptide-KLH complex in complete Freund's adjuvant. The resulting antisera are tested for antipeptide activity by binding the peptide to plastic, blocking with 1% bovine serum albumin (BSA), reacting with rabbit antisera, washing, and reacting with radioiodinated goat anti-rabbit IgG. Antisera with antipeptide activity are tested for anti-MDDT activity using protocols well known in the art, including ELISA, radioimmunoassay (RIA), and immunoblotting.

In another procedure, isolated and purified peptide may be used to immunize mice (about 100 µg of peptide) or rabbits (about 1 mg of peptide). Subsequently, the peptide is radioiodinated and used to screen the immunized animals' B-lymphocytes for production of antipeptide antibodies. Positive cells are then used to produce hybridomas using standard techniques. About 20 mg of peptide is sufficient for labeling and screening several thousand clones. Hybridomas of interest are detected by screening with radioiodinated peptide to identify those fusions producing peptide-specific monoclonal antibody. In a typical protocol, wells of a multi-well plate (FAST, Becton-Dickinson, Palo Alto, CA) are coated with affinity-purified, specific rabbit-anti-mouse (or suitable anti-species IgG) antibodies at 10 mg/ml. The coated wells are blocked with 1% BSA and washed and exposed to supernatants from hybridomas. After incubation, the wells are exposed to radiolabeled peptide at 1 mg/ml.

Clones producing antibodies bind a quantity of labeled peptide that is detectable above background. Such clones are expanded and subjected to 2 cycles of cloning. Cloned hybridomas are injected into pristane-treated mice to produce ascites, and monoclonal antibody is purified from the ascitic fluid by affinity chromatography on protein A (Amersham Pharmacia Biotech). Several procedures for the production of monoclonal antibodies, including in vitro production, are described in Pound (supra). Monoclonal antibodies with antipeptide activity are tested for anti-MDDT activity using protocols well known in the art, including ELISA, RIA, and immunoblotting.

Antibody fragments containing specific binding sites for an epitope may also be generated. For example, such fragments include, but are not limited to, the F(ab')2 fragments produced by pepsin digestion of the antibody molecule, and the Fab fragments generated by reducing the disulfide bridges of the F(ab')2 fragments. Alternatively, construction of Fab expression libraries in filamentous bacteriophage allows rapid and easy identification of monoclonal fragments with desired specificity (Pound, supra, Chaps. 45-47). Antibodies generated against polypeptide encoded by mddt can be used to purify and characterize full-length MDDT protein and its activity, binding partners, etc.

Assays Using Antibodies

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Anti-MDDT antibodies may be used in assays to quantify the amount of MDDT found in a particular human cell. Such assays include methods utilizing the antibody and a label to detect expression level under normal or disease conditions. The peptides and antibodies of the invention may be used with or without modification or labeled by joining them, either covalently or noncovalently, with a reporter molecule.

Protocols for detecting and measuring protein expression using either polyclonal or monoclonal antibodies are well known in the art. Examples include ELISA, RIA, and fluorescent activated cell sorting (FACS). Such immunoassays typically involve the formation of complexes between the MDDT and its specific antibody and the measurement of such complexes. These and other assays are described in Pound (supra).

Without further elaboration, it is believed that one skilled in the art can, using the preceding description, utilize the present invention to its fullest extent. The following preferred specific embodiments are, therefore, to be construed as merely illustrative, and not limitative of the remainder of the disclosure in any way whatsoever.

The disclosures of all patents, applications and publications, mentioned above and below, including U.S. Ser. No. 60/349,946 and U.S. 60/349,413, are hereby expressly incorporated by reference.

EXAMPLES

20 I. Construction of cDNA Libraries

RNA was purchased from CLONTECH Laboratories, Inc. (Palo Alto CA) or isolated from various tissues. Some tissues were homogenized and lysed in guanidinium isothiocyanate, while others were homogenized and lysed in phenol or in a suitable mixture of denaturants, such as TRIZOL (Life Technologies), a monophasic solution of phenol and guanidine isothiocyanate. The resulting lysates were centrifuged over CsCl cushions or extracted with chloroform. RNA was precipitated with either isopropanol or sodium acetate and ethanol, or by other routine methods.

Phenol extraction and precipitation of RNA were repeated as necessary to increase RNA purity. In most cases, RNA was treated with DNase. For most libraries, poly(A+) RNA was isolated using oligo d(T)-coupled paramagnetic particles (Promega Corporation (Promega), Madison WI), OLIGOTEX latex particles (QIAGEN, Inc. (QIAGEN), Valencia CA), or an OLIGOTEX mRNA purification kit (QIAGEN). Alternatively, RNA was isolated directly from tissue lysates using other RNA isolation kits, e.g., the POLY(A)PURE mRNA purification kit (Ambion, Inc., Austin TX).

In some cases, Stratagene was provided with RNA and constructed the corresponding cDNA libraries. Otherwise, cDNA was synthesized and cDNA libraries were constructed with the UNIZAP vector system (Stratagene Cloning Systems, Inc. (Stratagene), La Jolla CA) or SUPERSCRIPT

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plasmid system (Life Technologies), using the recommended procedures or similar methods known in the art. (See, e.g., Ausubel, 1997, supra, Chapters 5.1 through 6.6.) Reverse transcription was initiated using oligo d(T) or random primers. Synthetic oligonucleotide adapters were ligated to double stranded cDNA, and the cDNA was digested with the appropriate restriction enzyme or enzymes. For most libraries, the cDNA was size-selected (300-1000 bp) using SEPHACRYL S1000, SEPHAROSE CL2B, or SEPHAROSE CL4B column chromatography (Amersham Pharmacia Biotech) or preparative agarose gel electrophoresis. cDNAs were ligated into compatible restriction enzyme sites of the polylinker of a suitable plasmid, e.g., PBLUESCRIPT plasmid (Stratagene), PSPORT1 plasmid (Life Technologies), PCDNA2.1 plasmid (Invitrogen, Carlsbad CA), PBK-CMV plasmid (Stratagene), PCR2-TOPOTA plasmid (Invitrogen), PCMV-ICIS plasmid (Stratagene), pIGEN (Incyte Genomics, Palo Alto CA), pRARE (Incyte Genomics), or pINCY (Incyte Genomics), or derivatives thereof. Recombinant plasmids were transformed into competent E. coli cells including XL1-Blue, XL1-BlueMRF, or SOLR from Stratagene or DH5α, DH10B, or ElectroMAX DH10B from Life Technologies.

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II. Isolation of cDNA Clones

Plasmids were recovered from host cells by <u>in vivo</u> excision using the UNIZAP vector system (Stratagene) or by cell lysis. Plasmids were purified using at least one of the following: the Magic or WIZARD Minipreps DNA purification system (Promega); the AGTC Miniprep purification kit (Edge BioSystems, Gaithersburg MD); and the QIAWELL 8, QIAWELL 8 Plus, and QIAWELL 8 Ultra plasmid purification systems or the R.E.A.L. PREP 96 plasmid purification kit (QIAGEN). Following precipitation, plasmids were resuspended in 0.1 ml of distilled water and stored, with or without lyophilization, at 4°C.

Alternatively, plasmid DNA was amplified from host cell lysates using direct link PCR in a high-throughput format. (Rao, V.B. (1994) Anal. Biochem. 216:1-14.) Host cell lysis and thermal cycling steps were carried out in a single reaction mixture. Samples were processed and stored in 384-well plates, and the concentration of amplified plasmid DNA was quantified fluorometrically using PICOGREEN dye (Molecular Probes, Inc. (Molecular Probes), Eugene OR) and a FLUOROSKAN II fluorescence scanner (Labsystems Oy, Helsinki, Finland).

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III. Sequencing and Analysis

cDNA sequencing reactions were processed using standard methods or high-throughput instrumentation such as the ABI CATALYST 800 thermal cycler (Applied Biosystems) or the PTC-200 thermal cycler (MJ Research) in conjunction with the HYDRA microdispenser (Robbins Scientific Corp., Sunnyvale CA) or the MICROLAB 2200 liquid transfer system (Hamilton). cDNA

sequencing reactions were prepared using reagents provided by Amersham Pharmacia Biotech or supplied in ABI sequencing kits such as the ABI PRISM BIGDYE Terminator cycle sequencing ready reaction kit (Applied Biosystems). Electrophoretic separation of cDNA sequencing reactions and detection of labeled polynucleotides were carried out using the MEGABACE 1000 DNA sequencing system (Molecular Dynamics); the ABI PRISM 373 or 377 sequencing system (Applied Biosystems) in conjunction with standard ABI protocols and base calling software; or other sequence analysis systems known in the art. Reading frames within the cDNA sequences were identified using standard methods (reviewed in Ausubel, 1997, supra, Chapter 7.7). Some of the cDNA sequences were selected for extension using the techniques disclosed in Example VIII.

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IV. Assembly and Analysis of Sequences

Component sequences from chromatograms were subject to PHRED analysis and assigned a quality score. The sequences having at least a required quality score were subject to various preprocessing editing pathways to eliminate, e.g., low quality 3'ends, vector and linker sequences, polyA tails, Alu repeats, mitochondrial and ribosomal sequences, bacterial contamination sequences, and sequences smaller than 50 base pairs. In particular, low-information sequences and repetitive elements (e.g., dinucleotide repeats, Alu repeats, etc.) were replaced by "n's", or masked, to prevent spurious matches.

Processed sequences were then subject to assembly procedures in which the sequences were assigned to gene bins (bins). Each sequence could only belong to one bin. Sequences in each gene bin were assembled to produce consensus sequences (templates). Subsequent new sequences were added to existing bins using BLASTN (v.1.4 WashU) and CROSSMATCH. Candidate pairs were identified as all BLAST hits having a quality score greater than or equal to 150. Alignments of at least 82% local identity were accepted into the bin. The component sequences from each bin were assembled using a version of PHRAP. Bins with several overlapping component sequences were assembled using DEEP PHRAP. The orientation (sense or antisense) of each assembled template was determined based on the number and orientation of its component sequences. Template sequences as disclosed in the sequence listing correspond to sense strand sequences (the "forward" reading frames), to the best determination. The complementary (antisense) strands are inherently disclosed herein. The component sequences which were used to assemble each template consensus sequence are listed in Table 5, along with their positions along the template nucleotide sequences.

Bins were compared against each other and those having local similarity of at least 82% were combined and reassembled. Reassembled bins having templates of insufficient overlap (less than 95% local identity) were re-split. Assembled templates were also subject to analysis by STITCHER/EXON MAPPER algorithms which analyze the probabilities of the presence of splice

variants, alternatively spliced exons, splice junctions, differential expression of alternative spliced genes across tissue types or disease states, etc. These resulting bins were subject to several rounds of the above assembly procedures.

Once gene bins were generated based upon sequence alignments, bins were clone joined based upon clone information. If the 5' sequence of one clone was present in one bin and the 3' sequence from the same clone was present in a different bin, it was likely that the two bins actually belonged together in a single bin. The resulting combined bins underwent assembly procedures to regenerate the consensus sequences.

The final assembled templates were subsequently annotated using the following procedure. Template sequences were analyzed using BLASTN (v2.0, NCBI) versus gbpri (GenBank version 135). "Hits" were defined as an exact match having from 95% local identity over 200 base pairs through 100% local identity over 100 base pairs, or a homolog match having an E-value, i.e. a probability score, of $\leq 1 \times 10^{-8}$. The hits were subject to frameshift FASTx versus GENPEPT (GenBank version 135). (See Table 8). In this analysis, a homolog match was defined as having an E-value of $\leq 1 \times 10^{-8}$. The assembly method used above was described in "System and Methods for Analyzing Biomolecular Sequences," U.S.S.N. 09/276,534, filed March 25, 1999, and the LIFESEQ Gold user manual (Incyte) both incorporated by reference herein.

Following assembly, template sequences were subjected to motif, BLAST, and functional analyses, and categorized in protein hierarchies using methods described in, e.g., "Database System Employing Protein Function Hierarchies for Viewing Biomolecular Sequence Data," U.S. Patent Number 6,023,659; "Relational Database for Storing Biomolecular Information," U.S.S.N. 08/947,845, filed October 9, 1997; "Project-Based Full-Length Biomolecular Sequence Database," U.S. Patent Number 5,953,727; and "Relational Database and System for Storing Information Relating to Biomolecular Sequences," U.S.S.N. 09/034,807, filed March 4, 1998, all of which are incorporated by reference herein.

The template sequences were further analyzed by translating each template in all three forward reading frames and searching each translation against the Pfam database of hidden Markov model-based protein families and domains using the HMMER software package (available to the public from Washington University School of Medicine, St. Louis MO). Regions of templates which, when translated, contain similarity to Pfam consensus sequences are reported in Table 3, along with descriptions of Pfam protein domains and families. Only those Pfam hits with an E-value of $\leq 1 \times 10^{-3}$ are reported. (See also World Wide Web site http://pfam.wustl.edu/ for detailed descriptions of Pfam protein domains and families.)

Additionally, the template sequences were translated in all three forward reading frames, and each translation was searched against hidden Markov models for signal peptides using the HMMER

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software package. Construction of hidden Markov models and their usage in sequence analysis has been described. (See, for example, Eddy, S.R. (1996) Curr. Opin. Str. Biol. 6:361-365.) Only those signal peptide hits with a cutoff score of 11 bits or greater are reported. A cutoff score of 11 bits or greater corresponds to at least about 91-94% true-positives in signal peptide prediction. Template sequences were also translated in all three forward reading frames, and each translation was searched against TMHMMER, a program that uses a hidden Markov model (HMM) to delineate transmembrane segments on protein sequences and determine orientation (Sonnhammer, E.L. et al. (1998) Proc. Sixth Intl. Conf. On Intelligent Systems for Mol. Biol., Glasgow et al., eds., The Am. Assoc. for Artificial Intelligence (AAAI) Press, Menlo Park, CA, and MIT Press, Cambridge, MA, pp. 175-182.) Regions of templates which, when translated, contain similarity to signal peptide or transmembrane consensus sequences are reported in Table 4.

The results of HMMER analysis as reported in Tables 3 and 4 may support the results of BLAST analysis as reported in Table 2 or may suggest alternative or additional properties of template-encoded polypeptides not previously uncovered by BLAST or other analyses.

Template sequences are further analyzed using the bioinformatics tools listed in Table 8, or using sequence analysis software known in the art such as MACDNASIS PRO software (Hitachi Software Engineering, South San Francisco CA) and LASERGENE software (DNASTAR). Template sequences may be further queried against public databases such as the GenBank rodent, mammalian, vertebrate, prokaryote, and eukaryote databases.

The template sequences were translated to derive the corresponding longest open reading frame as presented by the polypeptide sequences as reported in Table 7. Alternatively, a polypeptide of the invention may begin at any of the methionine residues within the full length translated polypeptide. Polypeptide sequences were subsequently analyzed by querying against the GenBank protein database (GENPEPT, (GenBank version 135)). Full length polynucleotide sequences are also analyzed using MACDNASIS PRO software (Hitachi Software Engineering, South San Francisco CA) and LASERGENE software (DNASTAR). Polynucleotide and polypeptide sequence alignments are generated using default parameters specified by the CLUSTAL algorithm as incorporated into the MEGALIGN multisequence alignment program (DNASTAR), which also calculates the percent identity between aligned sequences.

Table 7 shows sequences with homology to the polypeptides of the invention as identified by BLAST analysis against the GenBank protein (GENPEPT) database. Column 1 shows the polypeptide sequence identification number (SEQ ID NO:) for the polypeptide segments of the invention. Column 2 shows the reading frame used in the translation of the polynucleotide sequences encoding the polypeptide segments. Column 3 shows the length of the translated polypeptide segments. Columns 4 and 5 show the start and stop nucleotide positions of the polynucleotide

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sequences encoding the polypeptide segments. Column 6 shows the GenBank identification number (GI Number) of the nearest GenBank homolog. Column 7 shows the probability score for the match between each polypeptide and its GenBank homolog. Column 8 shows the annotation of the GenBank homolog.

V. Analysis of Polynucleotide Expression

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Northern analysis is a laboratory technique used to detect the presence of a transcript of a gene and involves the hybridization of a labeled nucleotide sequence to a membrane on which RNAs from a particular cell type or tissue have been bound. (See, e.g., Sambrook, supra, ch. 7; Ausubel, 1995, supra, ch. 4 and 16.)

Analogous computer techniques applying BLAST were used to search for identical or related molecules in cDNA databases such as GenBank or LIFESEQ (Incyte Genomics). This analysis is much faster than multiple membrane-based hybridizations. In addition, the sensitivity of the computer search can be modified to determine whether any particular match is categorized as exact or similar. The basis of the search is the product score, which is defined as:

BLAST Score x Percent Identity

5 x minimum {length(Seq. 1), length(Seq. 2)}

The product score takes into account both the degree of similarity between two sequences and the length of the sequence match. The product score is a normalized value between 0 and 100, and is calculated as follows: the BLAST score is multiplied by the percent nucleotide identity and the product is divided by (5 times the length of the shorter of the two sequences). The BLAST score is calculated by assigning a score of +5 for every base that matches in a high-scoring segment pair (HSP), and -4 for every mismatch. Two sequences may share more than one HSP (separated by gaps). If there is more than one HSP, then the pair with the highest BLAST score is used to calculate the product score. The product score represents a balance between fractional overlap and quality in a BLAST alignment. For example, a product score of 100 is produced only for 100% identity over the entire length of the shorter of the two sequences being compared. A product score of 70 is produced either by 100% identity and 70% overlap at one end, or by 88% identity and 100% overlap at the other. A product score of 50 is produced either by 100% identity and 50% overlap at one end, or 79% identity and 100% overlap.

VI. Tissue Distribution Profiling

A tissue distribution profile is determined for each template by compiling the cDNA library tissue classifications of its component cDNA sequences. Each component sequence, is derived from

a cDNA library constructed from a human tissue. Each human tissue is classified into one of the following categories: cardiovascular system; connective tissue; digestive system; embryonic structures; endocrine system; exocrine glands; genitalia, female; genitalia, male; germ cells; hemic and immune system; liver; musculoskeletal system; nervous system; pancreas; respiratory system; sense organs; skin; stomatognathic system; unclassified/mixed; or urinary tract. Template sequences, component sequences, and cDNA library/tissue information are found in the LIFESEQ GOLD database (Incyte Genomics, Palo Alto CA).

Table 6 shows the tissue distribution profile for the templates of the invention. For each template, the three most frequently observed tissue categories are shown in column 3, along with the percentage of component sequences belonging to each category. Only tissue categories with percentage values of $\geq 10\%$ are shown. A tissue distribution of "widely distributed" in column 3 indicates percentage values of <10% in all tissue categories.

VII. Transcript Image Analysis

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Transcript images are generated as described in Seilhamer et al., "Comparative Gene Transcript Analysis," U.S. Patent Number 5,840,484, incorporated herein by reference.

VIII. Extension of Polynucleotide Sequences and Isolation of a Full-length cDNA

Oligonucleotide primers designed using an mddt of the Sequence Listing are used to extend the nucleic acid sequence. One primer is synthesized to initiate 5' extension of the template, and the other primer, to initiate 3' extension of the template. The initial primers may be designed using OLIGO 4.06 software (National Biosciences, Inc. (National Biosciences), Plymouth MN), or another appropriate program, to be about 22 to 30 nucleotides in length, to have a GC content of about 50% or more, and to anneal to the target sequence at temperatures of about 68°C to about 72°C. Any stretch of nucleotides which would result in hairpin structures and primer-primer dimerizations are avoided. Selected human cDNA libraries are used to extend the sequence. If more than one extension is necessary or desired, additional or nested sets of primers are designed.

High fidelity amplification is obtained by PCR using methods well known in the art. PCR is performed in 96-well plates using the PTC-200 thermal cycler (MJ Research). The reaction mix contains DNA template, 200 nmol of each primer, reaction buffer containing Mg²⁺, (NH₄)₂SO₄, and β-mercaptoethanol, Taq DNA polymerase (Amersham Pharmacia Biotech), ELONGASE enzyme (Life Technologies), and Pfu DNA polymerase (Stratagene), with the following parameters for primer pair PCI A and PCI B: Step 1: 94°C, 3 min; Step 2: 94°C, 15 sec; Step 3: 60°C, 1 min; Step 4: 68°C, 2 min; Step 5: Steps 2, 3, and 4 repeated 20 times; Step 6: 68°C, 5 min; Step 7: storage at 4°C. In the alternative, the parameters for primer pair T7 and SK+ are as follows: Step 1: 94°C, 3 min; Step 2:

94°C, 15 sec; Step 3: 57°C, 1 min; Step 4: 68°C, 2 min; Step 5: Steps 2, 3, and 4 repeated 20 times; Step 6: 68°C, 5 min; Step 7: storage at 4°C.

The concentration of DNA in each well is determined by dispensing 100 μ l PICOGREEN quantitation reagent (0.25% (v/v); Molecular Probes) dissolved in 1X Tris-EDTA (TE) and 0.5 μ l of undiluted PCR product into each well of an opaque fluorimeter plate (Corning Incorporated (Corning), Corning NY), allowing the DNA to bind to the reagent. The plate is scanned in a FLUOROSKAN II (Labsystems Oy) to measure the fluorescence of the sample and to quantify the concentration of DNA. A 5 μ l to 10 μ l aliquot of the reaction mixture is analyzed by electrophoresis on a 1% agarose mini-gel to determine which reactions are successful in extending the sequence.

The extended nucleotides are desalted and concentrated, transferred to 384-well plates, digested with CviJI cholera virus endonuclease (Molecular Biology Research, Madison WI), and sonicated or sheared prior to religation into pUC 18 vector (Amersham Pharmacia Biotech). For shotgun sequencing, the digested nucleotides are separated on low concentration (0.6 to 0.8%) agarose gels, fragments are excised, and agar digested with AGAR ACE (Promega). Extended clones are religated using T4 ligase (New England Biolabs, Inc., Beverly MA) into pUC 18 vector (Amersham Pharmacia Biotech), treated with Pfu DNA polymerase (Stratagene) to fill-in restriction site overhangs, and transfected into competent <u>E. coli</u> cells. Transformed cells are selected on antibiotic-containing media, individual colonies are picked and cultured overnight at 37°C in 384-well plates in LB/2x carbenicillin liquid media.

The cells are lysed, and DNA is amplified by PCR using Taq DNA polymerase (Amersham Pharmacia Biotech) and Pfu DNA polymerase (Stratagene) with the following parameters: Step 1: 94°C, 3 min; Step 2: 94°C, 15 sec; Step 3: 60°C, 1 min; Step 4: 72°C, 2 min; Step 5: steps 2, 3, and 4 repeated 29 times; Step 6: 72°C, 5 min; Step 7: storage at 4°C. DNA is quantified by PICOGREEN reagent (Molecular Probes) as described above. Samples with low DNA recoveries are reamplified using the same conditions as described above. Samples are diluted with 20% dimethysulfoxide (1:2, v/v), and sequenced using DYENAMIC energy transfer sequencing primers and the DYENAMIC DIRECT kit (Amersham Pharmacia Biotech) or the ABI PRISM BIGDYE Terminator cycle sequencing ready reaction kit (Applied Biosystems).

In like manner, the mddt is used to obtain regulatory sequences (promoters, introns, and enhancers) using the procedure above, oligonucleotides designed for such extension, and an appropriate genomic library.

IX. Labeling of Probes and Southern Hybridization Analyses

Hybridization probes derived from the mddt of the Sequence Listing are employed for screening cDNAs, mRNAs, or genomic DNA. The labeling of probe nucleotides between 100 and

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1000 nucleotides in length is specifically described, but essentially the same procedure may be used with larger cDNA fragments. Probe sequences are labeled at room temperature for 30 minutes using a T4 polynucleotide kinase, γ^{32} P-ATP, and 0.5X One-Phor-All Plus (Amersham Pharmacia Biotech) buffer and purified using a ProbeQuant G-50 Microcolumn (Amersham Pharmacia Biotech). The probe mixture is diluted to 10^7 dpm/ μ g/ml hybridization buffer and used in a typical membrane-based hybridization analysis.

The DNA is digested with a restriction endonuclease such as Eco RV and is electrophoresed through a 0.7% agarose gel. The DNA fragments are transferred from the agarose to nylon membrane (NYTRAN Plus, Schleicher & Schuell, Inc., Keene NH) using procedures specified by the manufacturer of the membrane. Prehybridization is carried out for three or more hours at 68°C, and hybridization is carried out overnight at 68°C. To remove non-specific signals, blots are sequentially washed at room temperature under increasingly stringent conditions, up to 0.1x saline sodium citrate (SSC) and 0.5% sodium dodecyl sulfate. After the blots are placed in a PHOSPHORIMAGER cassette (Molecular Dynamics) or are exposed to autoradiography film, hybridization patterns of standard and experimental lanes are compared. Essentially the same procedure is employed when screening RNA.

X. Chromosome Mapping of mddt

The cDNA sequences which were used to assemble SEQ ID NO:1-104 are compared with sequences from the Incyte LIFESEQ database and public domain databases using BLAST and other implementations of the Smith-Waterman algorithm. Sequences from these databases that match SEQ ID NO:1-104 are assembled into clusters of contiguous and overlapping sequences using assembly algorithms such as PHRAP (Table 8). Radiation hybrid and genetic mapping data available from public resources such as the Stanford Human Genome Center (SHGC), Whitehead Institute for Genome Research (WIGR), and Généthon are used to determine if any of the clustered sequences have been previously mapped. Inclusion of a mapped sequence in a cluster will result in the assignment of all sequences of that cluster, including its particular SEQ ID NO:, to that map location. The genetic map locations of SEQ ID NO:1-104 are described as ranges, or intervals, of human chromosomes. The map position of an interval, in centiMorgans, is measured relative to the terminus of the chromosome's p-arm. (The centiMorgan (cM) is a unit of measurement based on recombination frequencies between chromosomal markers. On average, 1 cM is roughly equivalent to 1 megabase (Mb) of DNA in humans, although this can vary widely due to hot and cold spots of recombination.) The cM distances are based on genetic markers mapped by Généthon which provide boundaries for radiation hybrid markers whose sequences were included in each of the clusters.

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XI. Microarray Analysis

Probe Preparation from Tissue or Cell Samples

Total RNA is isolated from tissue samples using the guanidinium thiocyanate method and polyA+RNA is purified using the oligo (dT) cellulose method. Each polyA+RNA sample is reverse transcribed using MMLV reverse-transcriptase, 0.05 pg/µl oligo-dT primer (21mer), 1X first strand buffer, 0.03 units/μl RNase inhibitor, 500 μM dATP, 500 μM dGTP, 500 μM dTTP, 40 μM dCTP, 40 μM dCTP-Cy3 (BDS) or dCTP-Cy5 (Amersham Pharmacia Biotech). The reverse transcription reaction is performed in a 25 ml volume containing 200 ng polyA+ RNA with GEMBRIGHT kits (Incyte). Specific control polyA+ RNAs are synthesized by in vitro transcription from non-coding yeast genomic DNA (W. Lei, unpublished). As quantitative controls, the control mRNAs at 0.002 ng, 0.02 ng, 0.2 ng, and 2 ng are diluted into reverse transcription reaction at ratios of 1:100,000, 1:10,000, 1:1000, 1:100 (w/w) to sample mRNA respectively. The control mRNAs are diluted into reverse transcription reaction at ratios of 1:3, 3:1, 1:10, 10:1, 1:25, 25:1 (w/w) to sample mRNA differential expression patterns. After incubation at 37°C for 2 hr, each reaction sample (one with Cy3 and another with Cy5 labeling) is treated with 2.5 ml of 0.5M sodium hydroxide and incubated for 20 minutes at 85°C to the stop the reaction and degrade the RNA. Probes are purified using two successive CHROMA SPIN 30 gel filtration spin columns (CLONTECH Laboratories, Inc. (CLONTECH), Palo Alto CA) and after combining, both reaction samples are ethanol precipitated using 1 ml of glycogen (1 mg/ml), 60 ml sodium acetate, and 300 ml of 100% ethanol. The probe is then dried to completion using a SpeedVAC (Savant Instruments Inc., Holbrook NY) and resuspended in 14 μ l 5X SSC/0.2% SDS.

Microarray Preparation

Sequences of the present invention are used to generate array elements. Each array element is amplified from bacterial cells containing vectors with cloned cDNA inserts. PCR amplification uses primers complementary to the vector sequences flanking the cDNA insert. Array elements are amplified in thirty cycles of PCR from an initial quantity of 1-2 ng to a final quantity greater than 5 μ g. Amplified array elements are then purified using SEPHACRYL-400 (Amersham Pharmacia Biotech).

Purified array elements are immobilized on polymer-coated glass slides. Glass microscope slides (Corning) are cleaned by ultrasound in 0.1% SDS and acetone, with extensive distilled water washes between and after treatments. Glass slides are etched in 4% hydrofluoric acid (VWR Scientific Products Corporation (VWR), West Chester, PA), washed extensively in distilled water, and coated with 0.05% aminopropyl silane (Sigma) in 95% ethanol. Coated slides are cured in a 110°C oven.

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Array elements are applied to the coated glass substrate using a procedure described in US Patent No. 5,807,522, incorporated herein by reference. 1 μ l of the array element DNA, at an average concentration of 100 ng/ μ l, is loaded into the open capillary printing element by a high-speed robotic apparatus. The apparatus then deposits about 5 nl of array element sample per slide.

Microarrays are UV-crosslinked using a STRATALINKER UV-crosslinker (Stratagene). Microarrays are washed at room temperature once in 0.2% SDS and three times in distilled water. Non-specific binding sites are blocked by incubation of microarrays in 0.2% casein in phosphate buffered saline (PBS) (Tropix, Inc., Bedford, MA) for 30 minutes at 60°C followed by washes in 0.2% SDS and distilled water as before.

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Hybridization

Hybridization reactions contain 9 μ l of probe mixture consisting of 0.2 μ g each of Cy3 and Cy5 labeled cDNA synthesis products in 5X SSC, 0.2% SDS hybridization buffer. The probe mixture is heated to 65° C for 5 minutes and is aliquoted onto the microarray surface and covered with an 1.8 cm² coverslip. The arrays are transferred to a waterproof chamber having a cavity just slightly larger than a microscope slide. The chamber is kept at 100% humidity internally by the addition of 140 μ l of 5x SSC in a corner of the chamber. The chamber containing the arrays is incubated for about 6.5 hours at 60° C. The arrays are washed for 10 min at 45° C in a first wash buffer (1X SSC, 0.1% SDS), three times for 10 minutes each at 45° C in a second wash buffer (0.1X SSC), and dried.

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<u>Detection</u>

Reporter-labeled hybridization complexes are detected with a microscope equipped with an Innova 70 mixed gas 10 W laser (Coherent, Inc., Santa Clara CA) capable of generating spectral lines at 488 nm for excitation of Cy3 and at 632 nm for excitation of Cy5. The excitation laser light is focused on the array using a 20X microscope objective (Nikon, Inc., Melville NY). The slide containing the array is placed on a computer-controlled X-Y stage on the microscope and raster-scanned past the objective. The 1.8 cm x 1.8 cm array used in the present example is scanned with a resolution of 20 micrometers.

In two separate scans, a mixed gas multiline laser excites the two fluorophores sequentially. Emitted light is split, based on wavelength, into two photomultiplier tube detectors (PMT R1477, Hamamatsu Photonics Systems, Bridgewater NJ) corresponding to the two fluorophores. Appropriate filters positioned between the array and the photomultiplier tubes are used to filter the signals. The emission maxima of the fluorophores used are 565 nm for Cy3 and 650 nm for Cy5. Each array is typically scanned twice, one scan per fluorophore using the appropriate filters at the laser source, although the apparatus is capable of recording the spectra from both fluorophores simultaneously.

The sensitivity of the scans is typically calibrated using the signal intensity generated by a cDNA control species added to the probe mix at a known concentration. A specific location on the array contains a complementary DNA sequence, allowing the intensity of the signal at that location to be correlated with a weight ratio of hybridizing species of 1:100,000. When two probes from different sources (e.g., representing test and control cells), each labeled with a different fluorophore, are hybridized to a single array for the purpose of identifying genes that are differentially expressed, the calibration is done by labeling samples of the calibrating cDNA with the two fluorophores and adding identical amounts of each to the hybridization mixture.

The output of the photomultiplier tube is digitized using a 12-bit RTI-835H analog-to-digital (A/D) conversion board (Analog Devices, Inc., Norwood, MA) installed in an IBM-compatible PC computer. The digitized data are displayed as an image where the signal intensity is mapped using a linear 20-color transformation to a pseudocolor scale ranging from blue (low signal) to red (high signal). The data is also analyzed quantitatively. Where two different fluorophores are excited and measured simultaneously, the data are first corrected for optical crosstalk (due to overlapping emission spectra) between the fluorophores using each fluorophore's emission spectrum.

A grid is superimposed over the fluorescence signal image such that the signal from each spot is centered in each element of the grid. The fluorescence signal within each element is then integrated to obtain a numerical value corresponding to the average intensity of the signal. The software used for signal analysis is the GEMTOOLS gene expression analysis program (Incyte Genomics). Array elements that exhibit at least about a two-fold change in expression, a signal-to-background ratio of at least about 2.5, and an element spot size of at least about 40%, are considered to be differentially expressed.).

XII. Complementary Nucleic Acids

Sequences complementary to the mddt are used to detect, decrease, or inhibit expression of the naturally occurring nucleotide. The use of oligonucleotides comprising from about 15 to 30 base pairs is typical in the art. However, smaller or larger sequence fragments can also be used. Appropriate oligonucleotides are designed from the mddt using OLIGO 4.06 software (National Biosciences) or other appropriate programs and are synthesized using methods standard in the art or ordered from a commercial supplier. To inhibit transcription, a complementary oligonucleotide is designed from the most unique 5' sequence and used to prevent transcription factor binding to the promoter sequence. To inhibit translation, a complementary oligonucleotide is designed to prevent ribosomal binding and processing of the transcript.

XIII. Expression of MDDT

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Expression and purification of MDDT is accomplished using bacterial or virus-based expression systems. For expression of MDDT in bacteria, cDNA is subcloned into an appropriate vector containing an antibiotic resistance gene and an inducible promoter that directs high levels of cDNA transcription. Examples of such promoters include, but are not limited to, the trp-lac (tac) hybrid promoter and the T5 or T7 bacteriophage promoter in conjunction with the lac operator regulatory element. Recombinant vectors are transformed into suitable bacterial hosts, e.g., BL21(DE3). Antibiotic resistant bacteria express MDDT upon induction with isopropyl beta-Dthiogalactopyranoside (IPTG). Expression of MDDT in eukaryotic cells is achieved by infecting insect or mammalian cell lines with recombinant Autographica californica nuclear polyhedrosis virus (AcMNPV), commonly known as baculovirus. The nonessential polyhedrin gene of baculovirus is replaced with cDNA encoding MDDT by either homologous recombination or bacterial-mediated transposition involving transfer plasmid intermediates. Viral infectivity is maintained and the strong polyhedrin promoter drives high levels of cDNA transcription. Recombinant baculovirus is used to infect Spodoptera frugiperda (Sf9) insect cells in most cases, or human hepatocytes, in some cases. Infection of the latter requires additional genetic modifications to baculovirus. (See e.g., Engelhard, supra; and Sandig, supra.)

In most expression systems, MDDT is synthesized as a fusion protein with, e.g., glutathione S-transferase (GST) or a peptide epitope tag, such as FLAG or 6-His, permitting rapid, single-step, affinity-based purification of recombinant fusion protein from crude cell lysates. GST, a 26-kilodalton enzyme from Schistosoma japonicum, enables the purification of fusion proteins on immobilized glutathione under conditions that maintain protein activity and antigenicity (Amersham Pharmacia Biotech). Following purification, the GST moiety can be proteolytically cleaved from MDDT at specifically engineered sites. FLAG, an 8-amino acid peptide, enables immunoaffinity purification using commercially available monoclonal and polyclonal anti-FLAG antibodies (Eastman Kodak Company, Rochester NY). 6-His, a stretch of six consecutive histidine residues, enables purification on metal-chelate resins (QIAGEN). Methods for protein expression and purification are discussed in Ausubel (1995, supra, Chapters 10 and 16). Purified MDDT obtained by these methods can be used directly in the following activity assay.

XIV. Demonstration of MDDT Activity

MDDT, or biologically active fragments thereof, are labeled with ¹²⁵I Bolton-Hunter reagent. (See, e.g., Bolton, A.E. and W.M. Hunter (1973) Biochem. J. 133:529-539.) Candidate molecules previously arrayed in the wells of a multi-well plate are incubated with the labeled MDDT, washed, and any wells with labeled MDDT complex are assayed. Data obtained using different

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concentrations of MDDT are used to calculate values for the number, affinity, and association of MDDT with the candidate molecules.

Alternatively, molecules interacting with MDDT are analyzed using the yeast two-hybrid system as described in Fields, S. and O. Song (1989) Nature 340:245-246, or using commercially available kits based on the two-hybrid system, such as the MATCHMAKER system (CLONTECH).

MDDT may also be used in the PATHCALLING process (CuraGen Corp., New Haven CT) which employs the yeast two-hybrid system in a high-throughput manner to determine all interactions between the proteins encoded by two large libraries of genes (Nandabalan, K. et al. (2000) U.S. Patent No. 6,057,101).

10 XV. Functional Assays

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MDDT function is assessed by expressing mddt at physiologically elevated levels in mammalian cell culture systems. cDNA is subcloned into a mammalian expression vector containing a strong promoter that drives high levels of cDNA expression. Vectors of choice include pCMV SPORT (Life Technologies) and pCR3.1 (Invitrogen Corporation, Carlsbad CA), both of which contain the cytomegalovirus promoter. 5-10- μ g of recombinant vector are transiently transfected into a human cell line, preferably of endothelial or hematopoietic origin, using either liposome formulations or electroporation. 1-2 μ g of an additional plasmid containing sequences encoding a marker protein are co-transfected.

Expression of a marker protein provides a means to distinguish transfected cells from nontransfected cells and is a reliable predictor of cDNA expression from the recombinant vector. Marker proteins of choice include, e.g., Green Fluorescent Protein (GFP; CLONTECH), CD64, or a CD64-GFP fusion protein. Flow cytometry (FCM), an automated laser optics-based technique, is used to identify transfected cells expressing GFP or CD64-GFP and to evaluate the apoptotic state of the cells and other cellular properties.

FCM detects and quantifies the uptake of fluorescent molecules that diagnose events preceding or coincident with cell death. These events include changes in nuclear DNA content as measured by staining of DNA with propidium iodide; changes in cell size and granularity as measured by forward light scatter and 90 degree side light scatter; down-regulation of DNA synthesis as measured by decrease in bromodeoxyuridine uptake; alterations in expression of cell surface and intracellular proteins as measured by reactivity with specific antibodies; and alterations in plasma membrane composition as measured by the binding of fluorescein-conjugated Annexin V protein to the cell surface. Methods in flow cytometry are discussed in Ormerod, M. G. (1994) Flow Cytometry, Oxford, New York NY.

The influence of MDDT on gene expression can be assessed using highly purified populations of cells transfected with sequences encoding MDDT and either CD64 or CD64-GFP.

CD64 and CD64-GFP are expressed on the surface of transfected cells and bind to conserved regions of human immunoglobulin G (IgG). Transfected cells are efficiently separated from nontransfected cells using magnetic beads coated with either human IgG or antibody against CD64 (DYNAL, Inc., Lake Success NY). mRNA can be purified from the cells using methods well known by those of skill in the art. Expression of mRNA encoding MDDT and other genes of interest can be analyzed by northern analysis or microarray techniques.

XVI. Production of Antibodies

MDDT substantially purified using polyacrylamide gel electrophoresis (PAGE; see, e.g., Harrington, M.G. (1990) Methods Enzymol. 182:488-495), or other purification techniques, is used to immunize rabbits and to produce antibodies using standard protocols.

Alternatively, the MDDT amino acid sequence is analyzed using LASERGENE software (DNASTAR) to determine regions of high immunogenicity, and a corresponding peptide is synthesized and used to raise antibodies by means known to those of skill in the art. Methods for selection of appropriate epitopes, such as those near the C-terminus or in hydrophilic regions are well described in the art. (See, e.g., Ausubel, 1995, supra, Chapter 11.)

Typically, peptides 15 residues in length are synthesized using an ABI 431A peptide synthesizer (Applied Biosystems) using fmoc-chemistry and coupled to KLH (Sigma) by reaction with N-maleimidobenzoyl-N-hydroxysuccinimide ester (MBS) to increase immunogenicity. (See, e.g., Ausubel, <u>supra.</u>) Rabbits are immunized with the peptide-KLH complex in complete Freund's adjuvant. Resulting antisera are tested for antipeptide activity by, for example, binding the peptide to plastic, blocking with 1% BSA, reacting with rabbit antisera, washing, and reacting with radio-iodinated goat anti-rabbit IgG. Antisera with antipeptide activity are tested for anti-MDDT activity using protocols well known in the art, including ELISA, RIA, and immunoblotting.

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XVII. Purification of Naturally Occurring MDDT Using Specific Antibodies

Naturally occurring or recombinant MDDT is substantially purified by immunoaffinity chromatography using antibodies specific for MDDT. An immunoaffinity column is constructed by covalently coupling anti-MDDT antibody to an activated chromatographic resin, such as CNBr-activated SEPHAROSE (Amersham Pharmacia Biotech). After the coupling, the resin is blocked and washed according to the manufacturer's instructions.

Media containing MDDT are passed over the immunoaffinity column, and the column is washed under conditions that allow the preferential absorbance of MDDT (e.g., high ionic strength buffers in the presence of detergent). The column is eluted under conditions that disrupt

antibody/MDDT binding (e.g., a buffer of pH 2 to pH 3, or a high concentration of a chaotrope, such as urea or thiocyanate ion), and MDDT is collected.

All publications and patents mentioned in the above specification are herein incorporated by reference. Various modifications and variations of the described method and system of the invention will be apparent to those skilled in the art without departing from the scope and spirit of the invention. Although the invention has been described in connection with specific preferred embodiments, it should be understood that the invention as claimed should not be unduly limited to such specific embodiments. Indeed, various modifications of the above-described modes for carrying out the invention which are obvious to those skilled in the field of molecular biology or related fields are intended to be within the scope of the following claims.

TABLE 1

SEQ ID NO:	101115:010 12	SEQ ID NO:	ORF ID
1	LG:1447398.9:2002JAN18	105	LG:1447398.9.orf2:2002JAN18
2	LG:201488.3:2002JAN18	106	LG:201488.3.orf3:2002JAN18
3	LG:288410.6:2002JAN18	107	LG:288410.6.orf2:2002JAN18
4	LG:7682817.1:2002JAN18	108	LG:7682817.1.orf3:2002JAN18
5	LG:7685059.6:2002JAN18	109	LG:7685059.6.orf2:2002JAN18
6	LG:7689671.1:2002JAN18	110	LG:7689671.1.orf2:2002JAN18
7	LG:7689684.1:2002JAN18	111	LG:7689684.1.orf2:2002JAN18
8	LG:7762669.1:2002JAN18	112	LG:7762669.1.orf1:2002JAN18
9	LG:965822.1:2002JAN18	113	LG:965822.1.orf1:2002JAN18
10	LG:006394.31:2002JAN18	114	LG:006394.31.orf2:2002JAN18
11	LG:018258.1:2002JAN18	115	LG:018258.1.orf2:2002JAN18
12	LG:027320.5:2002JAN18	116	LG:027320.5.orf3:2002JAN18
13	LG:057499.1:2002JAN18	117	LG:057499.1.orf1:2002JAN18
14	LG:065935.21:2002JAN18	118	LG:065935.21.orf2:2002JAN18
15	LG:071860.12:2002JAN18	119	LG:071860.12.orf3:2002JAN18
16	LG:087383.29:2002JAN18	120	LG:087383.29.orf2:2002JAN18
17	LG:098580.3:2002JAN18	121	LG:098580.3.orf2:2002JAN18
18	LG:1001879.1:2002JAN18	122	LG:1001879.1.orf1:2002JAN18
19	LG:1079456.4:2002JAN18	123	LG:1079456.4.orf3:2002JAN18
20	LG:1080598.9:2002JAN18	124	LG:1080598.9.orf2:2002JAN18
21	LG:1090358.10:2002JAN18	125	LG:1090358.10.orf1:2002JAN18
22	LG:1097492.2:2002JAN18	126	LG:1097492.2.orf2:2002JAN18
23	LG:1099945.26:2002JAN18	127	LG:1099945.26.orf2:2002JAN18
24	LG:110016.1:2002JAN18	128	LG:110016.1.orf1:2002JAN18
25	LG:1137613.10:2002JAN18	129	LG:1137613.10.orf2:2002JAN18
26	LG:118836.26:2002JAN18	130	LG:118836.26.orf2:2002JAN18
27	LG:1330261.32:2002JAN18	131	LG:1330261.32.orf1:2002JAN18
28	LG:1347461.28:2002JAN18		LG:1347461.28.orf3:2002JAN18
29	LG:1383494.16:2002JAN18		LG:1383494.16.orf3:2002JAN18
30	LG:1400155.1:2002JAN18	134	LG:1400155.1.orf2:2002JAN18
31	LG:1446621.1:2002JAN18	135	LG:1446621.1.orf3:2002JAN18
32	LG:144920.1:2002JAN18	136	LG:144920.1.orf1:2002JAN18
33	LG:1452619.1:2002JAN18	137	LG:1452619.1.orf2:2002JAN18
34	LG:1453417.6:2002JAN18	138	LG:1453417.6.orf1:2002JAN18
35	LG:148485.8:2002JAN18	139	LG:148485.8.orf1:2002JAN18
36	LG:1502670.1:2002JAN18	140	LG:1502670.1.orf2:2002JAN18
37	LG:206593.3:2002JAN18	141	LG:206593.3.orf2:2002JAN18
38	LG:228273.22:2002JAN18	142	LG:228273.22.orf1:2002JAN18
39	LG:228319.2:2002JAN18	143	LG:228319.2.orf1:2002JAN18
40	LG:229165.16:2002JAN18	144	LG:229165.16.orf2:2002JAN18
41	LG:230895.9:2002JAN18	145	LG:230895.9.orf1:2002JAN18
42	LG:233552.5:2002JAN18	146	LG:233552.5.orf1:2002JAN18
43	LG:234430.7:2002JAN18	147	LG:234430.7.orf3:2002JAN18
44	LG:236659.1:2002JAN18	148	LG:236659.1.orf3:2002JAN18
45	LG:236767.26:2002JAN18	149	LG:236767.26.orf2:2002JAN18
46	LG:237489.7:2002JAN18	150	LG:237489.7.orf1:2002JAN18
47	LG:238218.20:2002JAN18	151	LG:238218.20.orf1:2002JAN18
48	LG:239939.14:2002JAN18	152	LG:239939.14.orf3:2002JAN18
49	LG:242288.11:2002JAN18	153	LG:242288.11.orf1:2002JAN18
50	LG:242491.29:2002JAN18	154	LG:242491.29.orf1:2002JAN18

TABLE 1

		IABLE I	
EQ ID NO:	Template ID	SEQ ID NO:	ORF ID
1	LG:243488.41:2002JAN18	155	LG:243488.41.orf3:2002JAN18
2	LG:247792.18:2002JAN18	156	LG:247792.18.orf2:2002JAN18
3	LG:253193.17:2002JAN18	157	LG:253193.17.orf3:2002JAN18
4	LG:257088.20:2002JAN18	158	LG:257088.20.orf2:2002JAN18
	LG:265552.1:2002JAN18	159	LG:265552.1.orf2:2002JAN18
55	LG:275355.12:2002JAN18	160	LG:275355.12.orf1:2002JAN18
56	LG:280014.1:2002JAN18	161	LG:280014.1.orf1:2002JAN18
57	LG:299937.3:2002JAN18	162	LG:299937.3.orf3:2002JAN18
58	LG:311197.3:2002JAN18	163	LG:311197.3.orf3:2002JAN18
59	LG:321069.2:2002JAN18	164	LG:321069.2.orf1:2002JAN18
60	LG:330900.8:2002JAN18	165	LG:330900.8.orf2:2002JAN18
61	LG:330931.9:2002JAN18	166	LG:330931.9.orf1:2002JAN18
62	LG:330985.1:2002JAN18	167	LG:330985.1.orf3:2002JAN18
63	LG:332027.9:2002JAN18	168	LG:332027.9.orf3:2002JAN18
64	LG:335377.8:2002JAN18	169	LG:335377.8.orf2:2002JAN18
65	LG:337452.25:2002JAN18	170	LG:337452.25.orf3:2002JAN18
66	LG:340580.16:2002JAN18	171	LG:340580.16.orf2:2002JAN18
67	LG:350272.6:2002JAN18	172	LG:350272.6.orf2:2002JAN18
68	LG:397228.1:2002JAN18	173	LG:397228.1.orf1:2002JAN18
69	LG:401325.41:2002JAN18	174	LG:401325.41.orf2:2002JAN18
70	LG:402029.14:2002JAN18	175	LG:402029.14.orf3:2002JAN18
71	LG:407233.2:2002JAN18	176	LG:407233.2.orf3:2002JAN18
72	LG:407346.1:2002JAN18	177	LG:407346.1.orf3:2002JAN18
73	LG:407689.7:2002JAN18	178	LG:407689.7.orf3:2002JAN18
74	LG:40700.1:2002JAN18	179	LG:407700.1.orf2:2002JAN18
75	LG:410461.92:2002JAN18		LG:410461.92.orf3:2002JAN18
76	LG:411043.3:2002JAN18	181	LG:411043.3.orf2:2002JAN18
77	LG:438690.47:2002JAN1		LG:438690.47.orf1:2002JAN18
78	LG:444677.81:2002JAN1		LG:444677.81.orf1:2002JAN18
79	LG:457464.24:2002JAN1		LG:457464.24.orf3:2002JAN18
80	LG:7684793.15:2002JAN		LG:7684793.15.orf3:2002JAN18
81	LG:/084/93.10.20029/11		IG:7687485.1.orf1:2002JAN18
82	LG:7687485.1:2002JAN1		LG:7689661.4.orf2:2002JAN18
83	LG:7689661.4:2002JAN1		IG:7690373.1.orf1:2002JAN18
84	LG:7690373.1:2002JAN1		1G:7696560,1,orf3:2002JAN18
85	LG:7696560.1:2002JAN1		LG:7698190.26.orf3:2002JAN18
86	LG:7698190.26:2002JAN		IG:7763560.12.orf1:2002JAN18
87	LG:7763560.12:2002JAN	118 192	IG:7763587.20.orf2:2002JAN18
88	LG:7763587.20:2002JAN		I G:899263.10.orf2:2002JAN18
89	LG:899263.10:2002JAN		IG:977837.31.orf1:2002JAN18
90	LG:977837.31:2002JAN	18 195	LG:978560.13.orf2:2002JAN18
91	LG:978560.13:2002JAN		LG:979390.2.orf1:2002JAN18
92	LG:979390.2:2002JAN1		LG:983019.1.orf2:2002JAN18
93	LG:983019.1:2002JAN1		LG:997202.7.orf2:2002JAN18
94	LG:997202.7:2002JAN1		IG:998756,3.orf1:2002JAN18
95	LG:998756.3:2002JAN1		LG:103460.28.orf2:2002JAN18
96	LG:103460.28:2002JAN	118 200	LG:1501505.19.orf1:2002JAN18
97	LG:1501505.19:2002JA	N18 201	LG:233444.9.orf2:2002JAN18
98	LG:233444.9:2002JAN	18 202	LG:234824.7.orf2:2002JAN18
99	LG:234824.7:2002JAN	18 203	LG:235708.23.orf1:2002JAN18
100	LG:235708.23:2002JAN	V18 204	LG.2007 00.20(0)(1)

TABLE 1

SEQ ID NO:	Template ID	SEQ ID NO:	
	LG:236649.14:2002JAN18	205	LG:236649.14.orf1:2002JAN18
	LG:332474.7:2002JAN18	206	LG:332474.7.orf1:2002JAN18
	LG:335727.8:2002JAN18	207	LG:335727.8.orf2:2002JAN18
	LG:481983.1:2002JAN18		LG:481983.1.orf3:2002JAN18
104	LG:401903.1.20023/1110	1200	2014017007

Annotation	Homo sapiens cDNA FLJ33450 fls, clone BRAMY2000025.	LIM domains containing protein 1 (Mus musculus)	ALS2CR7 (Homo sapiens)	Homo saplens, clone MGC: 15950 IMAGE: 3338227, 11113, 3, 4	Homo sapiens cDNA FLJ34942 IIS, CIOI 18 1912/N 7507 507 107 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 11 21 2	Homo saplens, clone IMAGE:4846514, MIRNA, Dalling Cas.	Homo sapiens culva rusa4721 iiis, cistra fragasasasas moderately similar to	Homo sapiens cDNA FLJ38144 tis, clone DyOsizoco 777, magazinia, 727, 2100 ZINC FINGER PROTEIN 91.	Homo sapiens, Similar to zinc finger protein 331; zinc III 19el protein 433, 2010	IMAGE:3920900, mRNA.	Homo sapiens, clone IMAGE:4/48000, IIIRANA, Parina cas.	RIKEN CUIVA UOTUUOYAUT BOTTO (MICE 17586)	Widedoo lastications fringitie motif protein TRIM33 alpha (TRIM33) mRNA, complete	cds; alternatively spliced.	Homo sapiens, clone MGC:43081 IN/AGE:3237/21, 11111 Company Co	Homo sapiens, mago-ridaill (Prosperilla), minum complete cds. clone MGC:17367 IMAGE:3861094, mRNA, complete cds.	Homo sapiens mRNA; cDNA DKF2p43441130 (110111 cloth) Complete cds.	Homo sapiens caspase 12 variant alpha (CASP12/11118197, COMPSS)	unnamed protein product (Homo saplens)	Homo sapiens cDNA FLJ90585 fis, clone PLACE 1005757, 118111, 2011	Homo sapiens, Similar to hypothetical protein FLJ20079, clone MGC:45408 Homo sapiens, Similar to hypothetical protein FLJ20079, clone MGC:45408	IMAGE:5540009, mkina, complete cas.
Probability Score	000	4.00E-98	0.00	1.00E-97	0.00	8.00E-45	0.00	1.00E-85	00'0		0.00	1.00E-110	200	00:00	00'0	1.00E-123	00:00	00:00	0.00	0.00	00'0	
GI Number	017/8002	921/40992	g0377007	g18203798	g21750806	g22477160	g21750781	g21754721	G 1955442	1	g20987337	g19483967	g9967126	g12407440	g23273088	g17390472	g12053194	g20069117	C21886479	g22760915	g21618498	,
SEQ Template ID		2	LG:201488.3:20023AN16	100	1	G-7680671.1:2002JAN18		LG:7762669.1:2002JAN18	SINAL COOC. L COOT. C. C.		10 1G-006394.31:2002JAN18	LG:018258.1:2002JAN18		LG:057499.1:2002JAN18	14 C-065035.21:2002JAN18	LG:071860.12:2002JAN18	16 LG:087383.29:2002JAN18	17 LG:098580.3:2002JAN18	81MA10000.t occioes 6	18 [G:10018/9.1.2002JAN18	1030598.9:2002JAN18	70 [19.1000070:7:200070]

SFO	SEO Template ID	GI Number	Probability	Annotation
₽			Score	
S S Z	LG:1090358.10:2002JAN18	g21749606	0.00	Homo sapiens cDNA FLJ33955 fis, clone CTONG2018652, moderately similar to ZINC FINGER PROTEIN MFG-3.
5	10-1007402 2-2002 IAN18	o10047329	0.00	KIAA1626 protein (Homo sapiens)
7 8	l m	g6808024	0.00	Homo sapiens mRNA; cDNA DKFZp434J0428 (from clone DKFZp434J0428).
2 2	LG:110016.1:2002JAN18	g16552018	0.00	Homo saplens cDNA FLJ32015 fis, clone NTONG 1000052, weakly similar to
				KOTTUS NOTVEGICUS II IKINA IOI NAICH TOTATA A 2500 AGNE AGC: 17810
22	LG:1137613.10:2002JAN18	g14789616	1.00E-175	Homo sapiens, similar to kinen curva of zagostos, garro, como mana il MAGE:3891655, mRNA, complete cds.
2	G:118836.26:2002JAN18	g12667437	00'0	Homo sapiens NIR3 mRNA, complete cds.
2 2	m	g22761032	0:00	unnamed protein product (Homo sapiens)
200	IG:1347461.28:2002JAN18 g21755436	g21755436	00'0	Homo sapiens cDNA FLJ38725 fis, clone KIDNE2010265.
8	1G:1383494.16:2002JAN18 g16741	g16741479	0.00	Homo sapiens, CGI-100 protein, clone MGC:5366 IMAGE:3046999, MikinA,
)		complete cds.
66 66	LG:1400155.1:2002JAN18	g556220	0.00	Human NAD+-dependent succinate-semialdehyde denydrogendse (33ADH)
}				mRNA, 3' end.
31	LG:1446621.1:2002JAN18	g16306805	1,00E-105	Homo sapiens, zinc finger protein 43 (HIF6), cione MeC. 10361 IMAGE. 3007 040,
				mRNA, complete cds.
8	1G-144920 1-2002JAN18	g2689444	1.00E-141	ZNF134 (Homo saplens)
3	IG:1452619.1;2002JAN18	g9246972	0.00	Homo sapiens RNA-binding protein BRUNOL2 (BRUNOL2) MKINA, COMPIEIE CAS.
25	IG:1453417.6:2002JAN18	g1665821	0.00	Similar to D.melanogaster cadherin-related tumor suppressor (notifice suppressor)
35	LG:148485,8;2002JAN18	g21595467	0.00	Homo sapiens, clone MGC:40579 IMAGE:521/3/2, mkNA, complete cas.
38	LG:1502670.1:2002JAN18	g21755190	0.00	Homo sapiens cDNA FLJ38528 fis, clone HCHONZUUV442.
37	LG:206593,3:2002JAN18	g16551839	0.00	Homo sapiens cDNA FLJ31875 fis, clone NIZKP/UUZ45U, Weakiy sii iiidi 10 zii 10
-)		FINGER PROTEIN 84.
38	LG:228273.22:2002JAN18	g3327175	00'0	Homo sapiens mRNA for KIAA0681 protein, partial cas.
8	LG:228319.2:2002JAN18	g21752073	0.00	unnamed protein product (Homo sapiens)
8		g3970716	0.00	Homo sapiens mRNA for KEI protein.
41	LG:230895.9:2002JAN18	g15620894	0.00	Homo sapiens mistra for KIAA1916 projein, portion cos.
42	LG:233552.5:2002JAN18	g10047282	0.00	Homo sapiens misina for Kida 1004 projeiri, parijai cas.
	1			

SEQ	SEQ Template ID	GI Number	Probability	Agnotation
ΩŞ			Score	
£4	LG:234430.7:2002JAN18	g12653106	00:00	Homo sapiens, hypothetical protein dJ37E16.5, clone MGC:8472 IMAGE:2821743, mRNA, complete cds.
4	LG:236659.1:2002JAN18	g12083895	00'0	Homo sapiens polybromo-1 (PB1) mRNA, complete cds, alternatively spliced.
45	ω	g18916868	00'0	Homo sapiens mRNA for KIAA1978 protein.
8	•	g16588680	00:00	Homo sapiens anion transporter/exchanger-9 (SLC26A9) mkNA, complete
47	LG:238218.20:2002JAN18	g19116222	0.00	Homo sapiens, clone MGC:9833 IMAGE:3863491, mRNA, complete cas.
8		g21758739	00'0	Homo sapiens cDNA FLJ25797 fts, clone TST0/046.
49	LG:242288,11:2002JAN18	g4164385	00:00	PAK4 protein (Homo saplens)
22	LG:242491.29:2002JAN18	g22760074	0.00	Homo sapiens cDNA FLJ90076 fis, clone HEMBA1004444, moderately similar to
				GLYCOPROTEIN 25L PRECURSOR.
5	LG:243488.41:2002JAN18	g21542540	0.00	Homo sapiens, Similar to HTPAP protein, clone MGC:32924 IMAGE:526/610,
				mRNA, complete cds.
25	LG:247792.18:2002JAN18	g21739660	00'0	Homo sapiens mRNA; cDNA DKFZp434L1426 (from clone DKFZp434L1426).
53	LG:253193.17:2002JAN18	g20988139	00:00	Homo sapiens, E74-like factor 1 (ets domain transcription factor), clone
}	•)		MGC:40398 IMAGE:4385989, mRNA, complete cds.
5	LG:257088.20:2002JAN18	g20372683	0.00	euchromatic histone methyltransferase 1 (Homo sapiens)
RS RS		g18490501	1,00E-100	RIKEN cDNA 2010002A20 gene (Mus musculus)
25	ω	g17223623	0.00	Homo sapiens ATP-binding cassette A9 mRNA, complete cds.
27		g18088579	00.00	Homo sapiens, clone MGC:23949 IMAGE:4243903, mRNA, complete cds.
28		g10439753	0.00	Homo sapiens cDNA; FLJ23158 fis, clone LNG09623.
22	LG:311197.3:2002JAN18	g21750727	00.00	Homo sapiens cDNA FLJ34876 fis, clone NT2NE2015362, moderately similar to
	·			Mitogen inducible gene mig-2.
8	LG:321069.2:2002JAN18	g18677068	0.00	Homo sapiens cDNA FL/23877 fis, clone LNG13624.
2	LG:330900.8:2002JAN18	g18043698	0.00	PRO2000 protein (Homo sapiens)
62	LG:330931.9:2002JAN18	g16551908	00'0	Homo sapiens cDNA FLJ31930 fis, clone NT2RP7006162, weakly similar to ZINC
				FINGER PROJEIN MIFG-3.
63	LG:330985.1:2002JAN18	g1222522		Human placental folate transporter (hPOLI I) mkINA, complete cas.
2	LG:332027.9:2002JAN18	g21749635		Homo sapiens cDNA FLJ33979 fis, clone DFNES2U0437 I.
65	LG:335377.8:2002JAN18	g16550148	0.00	Homo sapiens cDNA FLJ30864 fis, clone FEBIXAZUU4U91, nigniy similar 10 HOMO sapiens DINC finaer profein terf mRNA

SFO	SEO Template ID	GI Number	Probability	Annotation
₽			Score	
Ö				MINOTONIA COLOR SOCIOLITATION
%	LG:337452.25:2002JAN18	g16549250		Homo sapiens CDNA FLUJUIU IIS, CIONE BINGTIA 1000 104.
67	i	g10434656	0.00	Homo sapiens cDNA FLJ 12900 fts, clone IN 1214P 2004321.
89	1	g22859174		hypothetical protein (Homo sapiens)
8	1	g15741221	1.00E-17	gene overexpressed in astrocytoma (Homo sapiens)
8	1	g4126475	0.00	BAP2-alpha protein (Homo sapiens)
-	1	g10437932	0.00	Homo sapiens cDNA: FL/21771 fts, clone COLF///9.
72		g20810035	0.00	Homo sapiens, Fc receptor-like protein 3, clone MGC.34000 IIVIAGE,4040737,
! 				mRNA, complete cds.
73	LG:407346.1:2002JAN18	g21753085	0.00	unnamed protein product (Homo sapiens)
7		g13365844	0.00	Macaca fascicularis brain cDNA clone: ACCE-19502, Tuli Inseli sequel ICE.
75		g14198271	0.00	Homo sapiens, clone MGC:5352 IMAGE:3048106, MKINA, COMPERCUS.
74	<u>س</u>	g16877725	0.00	Homo sapiens, likely ortholog of mouse g1-related zinc linger profell to clothe
₹)		MGC:15167 IMAGE:3535930, mRNA, complete cds.
12	LG:411043.3;2002JAN18	g21755898	00:00	unnamed protein product (Homo sapiens)
2/2	1G:438690.47:2002JAN18	g10434222	00'0	Homo sapiens cDNA FLJ12622 fis, clone NLZKM4001/31, nigniy sirniiai 10 norra
<u>-</u>)		sapiens F-box protein Lilina (LILINA) mRNA.
2	LG:444677.81:2002JAN18	g4929678	00'0	Homo sapiens CGI-105 protein mRNA, complete cds.
8	LG:457464.24:2002JAN18	g12653266	00'0	Homo saplens, fusion, derived from f(12,16) malignant liposarcoma, clone
}		1		MGC:8537 IMAGE:2822692, mRNA, complete cds.
2	IG:7684793.15:2002JAN18	g12275895	00:00	Irripartite motif protein TRIM19 gamma (Homo sapiens)
8	LG:7687485.1:2002JAN18	g21752508	0.00	Homo sapiens cDNA FLJ36280 ffs, clone THYMU2UU3282, moderately sittilial to
				ZINC FINGER PROJEIN 135.
88	LG:7689661.4:2002JAN18	g21739829	00'0	Homo sapiens mRNA; cDNA DKFZp761C148 (from clone DKF2p701C149),
				complete cds.
8	1G:7690373.1:2002JAN18	g21757678	00'0	Homo sapiens cDNA FLJ40479 ffs, clone TES112043282, moderately similar 10
)		ZINC FINGER PROTEIN 43.
85	1G:7696560.1:2002JAN18	g21752388	00.00	Homo sapiens cDNA FLJ36178 fis, clone TEST12026534.
8	1	g4914583	0.00	Homo saplens mRNA; cDNA DKFZp586A032 (from clone DKF2p580AU32);
				partial cds.
87	LG:7763560.12:2002JAN18 g12005677	g12005677	0.00	Homo sapiens HT029 mRNA, complete cds.

_					_	Т	_	\neg			Т	Т	1	Т			T	T	T	\top					
	Annotation	MGC:3528	Homo sapiens, ankyrin repear-collidii iii le proteii y collidii	IMAGE:3607648, mRNA, complete cas.	Homo sapiens cDNA FLJ35U84 118, civille FLACECOCCATOR.	Homo sapiens cDNA FLI90440 118, ciolie inizia 3000721, 1100011.	TUMORY SUPPRESSOR PROTEIN BOOK INC.	KIAA1588 protein (Holling supremis)	Homo sapiens culva Fludyzaa iis, ciaria bomolog (Xenopus Idevis), clane	Homo sapiens, Similar to eomesodell III 1110 I 100 Scott of Table 1	IMAGE:5223017, mRNA.	Homo sapiens, clone IMAGE:3638276, mkinA, parrial cus.	NEW Jomain containing protein isoform (Homo saplens)	THEW I CONTROLL STATE CLONE OVARC1000466.	Homo sapletis culto italogo monto narial cds.	Homo sapiens, clone IIVIAGE, 3000740, 11111 V. P. C.	Innamed protein product (Homo sapiens)	Hamp scripers denomic sequence surrounding Noti site, clone INLI-CLIUM.	Homo scalens CDNA FL 190562 fis, clone OVARC1001163.	Holling supremised the control of the convenient	Durding leganing: P138969 fls. clone NT2Ri2002359.	HOLITO SUCIETA CENTRAL 125361 fls. clone 13101713.	Horring supriens clone IMAGE:4396549, mRNA, partial cds.	HOLLO SUDICIPIENT CONTRACTOR	
	Probability Annotation Score		0.00		0.00	00'0		0:00	1.00E-180	0.00		000	000	0.00	000	0.00	000	8 6	0.00	0.00	1.001-118	000	0.00	0.00	
	GI Number		g12803700		g21750988	g22760683		g10047251	q21756044	093334960	2000	700000	g12803934	g24181969	g7023281	715070456	3000134	9/023130	g158/9022	g22760880	g20067239	g21755742	g16554117	g21410797	
	SEQ Template ID		012803700 A12803700	1.0.1/1.0000/1.50.2002.02:	10:0000 10:000 IAN18			1 C-078560 13:2002JAN18	10:070300 0:0001 IAN18	LG:9/9390.2.20023/1100	[G:983019.1:20023A19			IG:998756.3:2002JAN18	10:103/60 28:2002 JAN18	LO. 100400: 10:0000 IANI 8	[G:1501505.17.2002.021.5]	LG:233444.9:2002JAN18	99 LG:234824.7:2002JAN18	100 LG:235708.23:2002JAN18	101 LG:236649.14:2002JAN18	102 LG:332474.7:2002JAN18	103 LG:335727.8:2002JAN18	104 1 C:481083 1:2002JAN18	1.0.401.207.104.01
	SEQ	<u>ک</u>	2 8	8	6	3 8		5	5 8	3/3	£		94	ક	2 2	3	6	8 9	8	8	[0	102	193	2	
																	O	ソ							

TABLE

Template ID	Start	Stop	Frame	Ptam HIT		
				1100	DDCI repeat	7.40E-14
LG:1447398.9:2002JAN18	677	754	torward 2	KPEL.	IM domain	2.10E-29
LG:201488.3:2002JAN18	57	233	forward 3	LIIVI	Protein kingse domain	7.40E-05
LG:288410.6:2002JAN18	443	1099	forward 2	pkindse	Protein kindse domain	3.70E-09
LG:288410.6;2002JAN18	243	1071	forward 1	Pilipsomal S5	Ribosomal protein S5, N-terminal domain	0.00051
LG:/08281/.1;2002JAN10	317	436	١.,	KRAB	KRAB box	1.90E-24
LG:/0000099.0.200207110	317	430		KRAB	KRAB box	1.90E-20
C:768068/ 1:7002.JAN18	188	310	forward 2	KRAB	KRAB box	8.20E-24
IG:7762669,1;2002JAN18	178	300	forward 1	KRAB	KRAB box	4.50E-25
IG:965822.1.2002JAN18	424	546	forward 1	KRAB	KRAB box	6 10F-57
LG:006394.31:2002JAN18	1790	2248	forward 2	RhoGAP	RhoGAP domain	3 20F-05
LG:006394.31:2002JAN18	228	317	forward 3	WW	WW domain	9.50E-05
LG:018258.1:2002JAN18	299	718	forward 2	Nitroreductase	Nitroreduciose talling	6,30E-06
LG:027320.5:2002JAN18	132	332	forward 3	SAM	SAM dolitidin (Sieine dipira mon)	4.10E-18
LG:057499.1:2002JAN18	1306	1575	forward 1	bromodomain	Bromodofituili	2.70E-12
LG:057499.1:2002JAN18	1087	1224	forward 1	몺	PHU-tinger	3.50E-36
LG:065935.21:2002JAN18	209	517	forward 2	HesB-IIke	Hesb-like domain	2 ANF-19
LG:071860.12:2002JAN18	426	773	forward 3	Mago_nashi	Mago nashi protein	2 ODF-30
LG:087383.29:2002JAN18	173	439	forward 2	준	Fes/CiP4 nomology dolition	1.30E-31
LG:098580.3:2002JAN18	74	331	forward 2	ICE p10	ICE-IIKe Dioledae (caspase) pro contrain	1 ADF-40
LG:1001879.1:2002JAN18	310	381	forward 1	LRR	Leucine Rich Repeat	4.40E-07
LG:1001879.1:2002JAN18	217	306	forward 1	LRRNT	Leucine rich repedi Iv-leit III Idi dori din	7.20E-05
LG:1001879.1:2002JAN18	1204	1365	forward 1	LRRCT	Leucine fichi lepedi Cieni ili di donicii i	1.00E-28
LG:1079456.4:2002JAN18	398	220	forward 2	KRAB	KIKAB DOX	4.00F-24
LG:1079456.4:2002JAN18	183	305	forward 3		_ [`	8,00E-59
LG:1080598.9:2002JAN18	755	823	forward 2	- 1	ZINC IIIIgei, Ozniz Iybe	1 80E-22
LG:1080598.9:2002JAN18	524	646	forward 2		KIKAB DOX	3 90F-27
LG:1090358.10:2002JAN18	3 808	88	forward 1	KRAB	KIKAB DOX	1 20F-22
LG:1090358.10:2002JAN18	3 1285	1353	forward 1	zf-C2H2	Zinc finger, CZHZ 190e	5 80F-52
LG:1090358.10:2002JAN18	3 1452	1520	forward 3	zf-C2H2	ZINC finger, CZHZ IYPE	2 OOF-05
RINAL CONCOUNTY	701	1107	forward 1		RhoGEL domain	200L 00

TABLE 3

E-value	1.80E-06	1.30E-23	8.40E-35	7.40E-20	1.10E-42	4.20E-11	1.30E-06	6.90E-07	9.90E-15	1.90E-115	4.00E-23	3.60E-71	2.90E-50		1.10E-76	4.00E-57	4.30E-31	1.50E-27	4.60E-18	1.60E-17	2.20E-81	8.10E-27	2.70E-45		1.10E-25		1.10E-10	1.60E-11	1.00E-46	2.00E-21	1 001 100
Pfam Description	Ribosomal protein L37e	Kelch motif	BTB/POZ domain	Dynein light chain type 1	DDHD domain	Calponin family repeat	Calponin homology (CH) domain	Uncharacterized ACR, YfiH family COG1496	emp24/gp25L/p24 family	Aldehyde dehydrogenase family	KRAB box	Zinc finger, C2H2 type	RNA recognition motif. (a.k.a. RRM, RBD, or RNP	domain)	Cadherin domain	7 transmembrane receptor (Secretin family)	EGF-like domain	Latrophilin/CL-1-like GPS domain	1	Hormone receptor domain	Cadherin domain	Cadherin domain	FGGY family of carbohydrate kinases, N-terminal	domain	FGGY family of carbohydrate kinases, C-terminal	domain	ADP-ribosylation factor family	Zinc finger, C2H2 type	Zinc finger, C2H2 type	Zinc finger, C2H2 type	the second field and
Pfam Hit	Ribosomal_L37e	Kelch	BTB	Dynein_light	DHQQ	calponin	당	DUF152	EMP24_GP25L	aldedh	KRAB	zf-C2H2	ırm		cadherin	7tm_2	EGF	GPS	laminin_G	HRM	cadherin	cadherin	FGGY		FGGY_C		arf	zf-C2H2	zf-C2H2	zf-C2H2	7900
Frame	forward 2	forward 2	forward 3	forward 2	forward 1	forward 3	forward 3		forward 3	forward 2	forward 3	forward 1	١.,		forward 1	forward 1	forward 1	forward 1	forward 1	forward 1	forward 2	forward 3	forward 1		forward 1		forward 2	forward 1	forward 2	1 1	I posorracy
Stop	364	1477	452	415	1770	5615	5441	1385	890	1408	668	252	403		2643		4080	7329	4896	6315		1064	852		1551		568	375		122	
Start	236	1337	135	152	1084	5538	5109	864	219	164	777	184	179		2374	7342	3919	7168	4408	6142	1436	780	49		859		2	307	476	54	627
Template ID	LG:1099945.26:2002JAN18	LG:110016.1:2002JAN18	LG:110016.1:2002JAN18	LG:1137613.10:2002JAN18	LG:118836.26:2002JAN18	LG:1330261.32:2002JAN18	LG:1330261.32:2002JAN18	LG:1347461.28:2002JAN18	LG:1383494,16:2002JAN18	LG:1400155.1:2002JAN18	LG:1446621.1:2002JAN18	LG:144920.1:2002JAN18	LG:1452619.1:2002JAN18		LG:1453417.6:2002JAN18	LG:1453417.6:2002JAN18	LG:1453417.6:2002JAN18	LG:1453417.6:2002JAN18	LG:1453417.6:2002JAN18	LG:1453417.6:2002JAN18	LG:1453417.6:2002JAN18	LG:1453417.6:2002JAN18	LG:148485.8:2002JAN18		LG:148485.8:2002JAN18		LG:1502670.1:2002JAN18	LG:206593.3:2002JAN18	LG:206593.3:2002JAN18	LG:206593.3:2002JAN18	9 LIN VI GUUG-GG 62 GGGG-C-1
SEQ	 23	24	24	25	56	27	27	28	29	တ္တ	31	32	33		34			34	34		34	34			35		36	37	37	37	

TABLE 3

i		_					
D NO:	011481 0000 00 0000	100	1004		JHC-J#	Zinc finger, C2HC type	3.40E-11
8	LG:228273.22:2002JAN18	100	000	for iord 1		SAM domain (Sterile alpha motif)	7.70E-08
88	LG:2282/3.22:2002JAN18	2194	1054	- 1		mbtrepeat	9.50E-81
65	LG:228319.2:2002JAN18	000	1670	\neg	100 H	mbtrepeat	1.60E-38
65	LG:228319.2:2002JAN18	155	1465	forward 2		P53	4.00E-196
3 5	LG:229100.10.2002301110	73	697	Т	Glycos transf 2	Glycosyl transferase	5.60E-37
4 5	1 G-230895.7.200237(118	1012	1137	forward 1	Ricin_B_lectin	QXW lectin repeat	7.70E-35
5 2	1G:233552.5:2002JAN18	874	1425		MIF4G	MIF4G domain	4.80E-3/
42	LG:233552,5:2002JAN18	1750	2070	forward 1	MA3	MA3 domain	1.90E-23
1 2	LG:234430,7:2002JAN18	123	848	forward 3	Hydrolase	haloacid dehalogenase-like hydrolase	1.0UE-21
2 4	IG:236659.1:2002JAN18	300	269	forward 3	bromodomain	Bromodomain	0.5UE-102
4	LG:236659.1:2002JAN18	3018	3374	1 1	BAH	BAH domain	0.00E-8/
4	IG-236659 1-2002JAN18	4290	4463	forward 3	HMG_box	HMG (high mobility group) box	0.00015
8	LG:236767.26:2002JAN18	197	391	forward 2	נגנט	RNA recognition motif. (a.k.a. RRM, RBD, or KNP)	I.10E-20
			100	F 44.00	CTAC	STAS domain	2.00E-20
8	LG:237489.7:2002JAN18	424	20.5	Forward 1	SAIS	KDAB hox	3.70E-14
47	LG:238218.20:2002JAN18	200	3/2	DI MOIO	OEND-B	CENP-8 protein	0.00056
47	LG:238218.20:2002JAN18	077	27.7	- pinwioi	בואו ה	Postfamily	1,90E-30
8	LG:239939.14:2002JAN18	3/8	6/0	S DIWOI	TOS Coscilla	Drotein kings domain	1.10E-62
4	LG:242288.11:2002JAN18	3	450 500	rorwara -	DKII TOSE	CHOISING CONTRAIN	3.00E-33
22	LG:242491.29:2002JAN18	2	663	forward	EMP24_GP25L	BADS amorphism (V	1.60E-32
51	LG:243488.41:2002JAN18	198	440	forward 3	PAPZ	PATE SUPERIOR IN	2,40E-115
52	LG:247792.18:2002JAN18	269	11/1	forward 2	KINB	KIND-IIKE DIOJEILI	1.00F-51
က္သ	LG:253193.17:2002JAN18	1854	2111	forward 3	Efs	EIS-doilidii	7 50F-32
54	LG:257088.20:2002JAN18	3230	3328	forward 2	ank	Ankyrin repedi	2 30F-09
<u>'¥</u>	LG:257088.20:2002JAN18	3531	3629	forward 3	ank	Ankyrin repedi	7 ODE-10
55	LG:265552.1:2002JAN18	242	427	forward 2	<u>D</u>	immunoglobulin doirrain	1 505-1
8	LG:275355.12:2002JAN18	49	468	forward 1	ABC_tran	ABC fransporrer	1 10E-16
57	LG:280014.1:2002JAN18	481	1005	forward 1	PMP22 Claudin	PMP-22/EMP/WP20/Cidudin Idiniiy	2 50E-06
88	LG:299937.3:2002JAN18	405	518	forward 3	zf-C3HC4	Zinc tinger, C3HC4 Type (Killy Silly Ber)	0.00055
28	11 G-200937 3-2002 JAN18	1008	1118	forward 3	zf-B box	B-box zinc tinger	00000

E-value	1.40E-09	7.70E-30	1.10E-80	7.60E-15	2.60E-39	5.00E-292	1.00E-90	2.50E-15	7.60E-12	5.60E-18	2.80E-10	6.10E-31	1.20E-17	1.10E-10	1.10E-11	0.00011	4.00E-40	1.80E-20	5.30E-27	3.00E-28	3.60E-19	2.10E-08	2.50E-20	1.30E-05	1.10E-19	1.20E-15	2.20E-08	1.30E-29	
Pfam Description	PH domain	Mitochondrial carrier protein	ATPase family associated with various cellular activities (AAA)	Bromodomain	Zinc finger, C2H2 type	Reduced folate carrier	MraW methylase family	B-box zinc finger	Zinc finger, C3HC4 type (RING finger)	SPRY domain	Protein kinase domain	PBX domain	Homeobox domain	SPRY domain	Zinc finger, C3HC4 type (RING finger)	SH3 domain	C Cation transporting ATPase, C-terminus	Immunoglobulin domain	Calponin homology (CH) domain	Leucine Rich Repeat	Protein phosphatase 2C	Phosphoglycerate mutase family	PA domain	Zinc finger, C3HC4 type (RING finger)	Zinc-binding dehydrogenase	CXXC zinc finger	F-box domain	Fumarylacetoacetate (FAA) hydrolase family	
Pfam Hit	H	mito_carr	AAA	bromodomain	zf-C2H2	Folate_carrier	Methyttransf_5	zf-B_box	zf-C3HC4	SPRY	pkinase	PBX	homeobox	SPRY	zf-C3HC4	SH3	Cation_ATPase_C	ğ	H)	LRR	PP2C	PGAM	PA	zf-C3HC4	adh_zinc	zf-CXXC	F-box	FAA_hydrolase	
Frame	forward 3	1	forward 2	forward 2	T -	_	1				forward 3				T		forward 3	I	forward 3		forward 3	forward 2	forward 3	!	forward 2	forward 1	forward 2	1	
Stop	1808	7	7	2821			1286	Ī		2033	1	02	Γ						602				764			069	П	1092	Ţ
Start	1497	544	965	2546	-	186	303	476	242	1671	3	1049	1643	947	49	1260	879	366			1191	338	468	993	287	550	1526	652	
Template ID	(G:311197.3:2002.IAN18	LG:321069.2:2002JAN18	LG:330900.8:2002JAN18	LG:330900.8:2002JAN18	LG:330931.9:2002JAN18	LG:330985,1:2002JAN18	LG:332027.9:2002JAN18	LG:335377.8:2002JAN18	LG:335377,8:2002JAN18	LG:335377.8:2002JAN18	LG:337452.25:2002JAN18	LG:340580.16:2002JAN18	LG:340580.16:2002JAN18	LG:350272.6:2002JAN18	LG:397228.1:2002JAN18	LG:401325.41:2002JAN18	LG:402029.14:2002JAN18	LG:407233.2:2002JAN18	LG:407346.1:2002JAN18	LG:407689.7:2002JAN18	LG:407689.7:2002JAN18	LG:407700.1:2002JAN18	LG:410461.92:2002JAN18	LG:410461.92:2002JAN18	LG:411043.3:2002JAN18	LG:438690.47:2002JAN18	LG:438690.47:2002JAN18	LG:444677.81:2002JAN18	
SEQ.	.50 S	9	61	61	62	63	49	65	65	92	99	67	67	89	69	2	71	72	73	74	74	75	7/6	76	77	78	78	79	

TABLE 3

RNA recognition motif. (a.k.a. RRM, RBD, or RNP domain) B-box zinc finger Zinc finger, C2H2 type Ankyrin repeat SAP domain ATP-NAD kinase Ankyrin repeat	zf-B box zf-C3HC4 zf-C3HC4 zf-C2H2 KRAB zf-C2H2 zf-C2H2 ank SAP NAD_kinase ank SAP NAD_kinase ank SAP NAD_kinase ank SAP SAP SAP SAP SAP SAP T-C2H2 ARAB T-C2H2 SAP SAP SAP SAP SAP SAP SAP SAP SAP SAP		4 9 4 8 2		0,000
ds.	8_box -C3HC4 -C2H2 -C2H2 -C2H2 -C2H2 -C2H2 -C2H2 -CC1 RAB RAB RAB RAB RAB RAB RAB RAB			forward 3 forward 2 forward 2 forward 2 forward 3 forward 3 forward 3 forward 3 forward 3 forward 2	384 512 forward 3 183 287 forward 3 46 114 forward 1 947 1006 forward 2 74 196 forward 2 684 752 forward 3 5 73 forward 3 5 73 forward 3 105 209 forward 3 714 1598 forward 3 714 1598 forward 3 714 1598 forward 2 455 598 forward 2 610 759 forward 2 610 759 forward 2 610 759 forward 2
dsi	F-C3HC4 F-C2H2 F-C2H2 F-C2H2 F-C2H2 F-C2H2 AP AP AND kindse AND ki			forward 3 forward 1 forward 2 forward 2 forward 3 forward 2 forward 3 forward 2 forward 3	183 287 forward 3 46 114 forward 1 947 1006 forward 2 74 196 forward 2 684 752 forward 3 5 73 forward 3 105 209 forward 3 714 1598 forward 3 714 1598 forward 3 714 1598 forward 3 715 2077 forward 2 455 598 forward 2 610 759 forward 2 610 759 forward 2 771 forward 2
Class Control of the	F-C2H2 RAB F-C2H2 F-C2H2 F-C2H2 AP AP AD_kinase ANCT CC1 RCC1 RRCT F-C2H2 RRCT RRCT RRCT F-C2H2 RRCT RRCT F-C2H2 RRCT RRCT F-C2H2 RRCT F-C2H2 RRCT RRCT RRCT RRCT F-C2H2 RRCT RRCT RRCT RRCT RRCT RRCT RRCT RRC	N N 저 N N 이 이 이 스 이 너 그 지 그	1 1 1 1 1 1 1 1 1 1 1 1	forward 1 forward 2 forward 2 forward 3 forward 3 forward 3 forward 3 forward 3 forward 3 forward 2 forward 2 forward 2 forward 2 forward 2	46 114 forward 1 947 1006 forward 2 74 196 forward 2 684 752 forward 3 5 73 forward 3 105 209 forward 3 714 1598 forward 3 714 1598 forward 3 610 759 forward 2
ds.	F-C2H2 F-C2H2 Ink		forward 2 forward 3 forward 3 forward 3 forward 3 forward 3 forward 2 forward 2 forward 1		947 1006 74 196 684 752 5 73 966 1064 105 209 714 1598 1979 2077 455 598 610 759 1115 1183
ds.	RAB F-C2H2 Ink AP VAD_kinase INK CC1 RRC1 RRC7 RRC8 RRAB (RAB RAB (RAB F-Dox		forward forwar		74 196 684 752 5 73 5 73 966 1064 105 209 714 1598 1979 2077 455 598 610 759 7115 1183
dsı	F-C2H2 Junk AP AAD kindse Junk ARD KINDSE ANK ARCT F-C2H2 CRAB CRAB CRAB F-box SH3 F-DHHC		forward forwar		684 752 5 73 966 1064 105 209 714 1598 1979 2077 455 598 610 759 1115 1183
ds			forward forward forward forward forward		5 73 966 1064 105 209 714 1598 1979 2077 455 598 610 759 1115 1183
dsı			forward forward forward forward		950 1004 105 209 714 1598 1979 2077 455 598 610 759 1115 1183
ds			forward forward forward forward		105 207 714 1598 1979 2077 455 598 610 759 1115 1183
ds	ank SCC1 RRCT rf-C2H2 (RAB (RAB (RAB (RAB (RAB (RAB (RAB (RAB		forward forward forward		1979 2077 455 598 610 759 1115 1183
	CCC1 RRCT FF-C2H2 (RAB CRAB F-box SH3 F-DHHC		forward		455 598 610 759 1115 1183
	rrct rf-C2H2 (RAB (RAB F-box 3H3		forward forward	1	759
	rf-C2H2 (RAB (RAB F-box 3H3 rf-DHHC		fond/ord		5 1183
	(RAB box box box 		2 2 2 2		COL
	(RAB 1-box 3H3 2f-DHHC		forward		
	-box 3H3 4FDHHC	Π.	forward		207
	ors P-DHHC		forward		220
	טייט-יי	\neg	torward		529
	<u> </u>		forward 2	1201 forward	1221
	Sulfate transp		forward 1		1 700
Domain of unknown function Leucine Rich Repeat	DUF300		forward		
Leucine Rich Repeat	DUF300	1	forward 2	Γ	3 946
togod dold original	LRR	1	forward 2		1150
Tencil le McII vebegi	182	\Box	forward 3		383
Domain of unknown function	DUF300		forward 3		2999
Zinc finger, C2H2 type	zf-C2H2	1	forward 1		2 456
KRAB box	KRAB		forward 2		469
WD domain, G-beta repeat	9	d2	forward 2 WD40		391

IABLE 3

							(: 10; · L
SEQ	Template ID	Start	Stop	Frame	Pfam Hilt	Pfam Description	E-value
ONO.							1 005 64
104	LG:481983.1:2002JAN18	414	953	forward 3	AG1	Longevity-assurance protein (LAG 1)	1.701-04

TABLE 4

				T.	·	
SEQ ID	Template ID	Start	Stop	Frame	l .	Topology
NO:					Туре	
10	LG:006394.31:2002JAN18	2653	2730	forward 1	SP	
10	LG:006394.31:2002JAN18	2653	2724	forward 1	SP	
10	LG:006394.31:2002JAN18	2653	2730	forward 1	SP	
10	LG:006394.31:2002JAN18	3539	3628	forward 2	SP	
10	LG:006394.31:2002JAN18	3539	3607	forward 2	SP	
10	LG:006394.31:2002JAN18	2874	2948	forward 3	SP	
11	LG:018258.1:2002JAN18	1	9		TM	Extracellular
11	LG:018258.1:2002JAN18	10	28		TM	Transmembrane
11	LG:018258.1:2002JAN18	29	278		TM	Cytosolic
11	LG:018258.1:2002JAN18	1	222		TM	Cytosolic
11	LG:018258.1:2002JAN18	223	245		TM	Transmembrane
11	LG:018258.1:2002JAN18	246	278		TM	Extracellular
11	LG:018258.1:2002JAN18	26	94	forward 2	SP	
11	LG:018258.1:2002JAN18	26	88	forward 2	SP	
11	LG:018258.1:2002JAN18	26	82	forward 2	SP	
11	LG:018258.1:2002JAN18	636	734	forward 3	SP	
11	LG:018258.1:2002JAN18	663	734	forward 3	SP	
 	LG:018258.1:2002JAN18	636	728	forward 3	SP	
11	LG:018258.1:2002JAN18	663	728	forward 3	SP	
11	LG:018258.1:2002JAN18	663	740	forward 3	SP	
11	LG:018258.1:2002JAN18	663	725	forward 3	SP	
11	LG:018258.1:2002JAN18	663	719	forward 3	SP	
11	LG:018258.1:2002JAN18	663	740	forward 3	SP	
11	LG:018258.1:2002JAN18	663	722	forward 3	SP	
12	LG:027320.5:2002JAN18	1	37		TM	Cytosolic
12	LG:027320.5:2002JAN18	38	60		TM	Transmembrane
12	LG:027320.5:2002JAN18	61	705		TM	Extracellular
12	LG:027320.5:2002JAN18	706	728		TM	Transmembrane
12	LG:027320.5:2002JAN18	729	801		TM	Cytosolic
12	LG:027320.5:2002JAN18	802	824		TM	Transmembrane
12	LG:027320.5:2002JAN18	825	843		TM	Extracellular
12	LG:027320.5:2002JAN18	844	866		TM	Transmembrane
12	LG:027320.5:2002JAN18	867	913		TM	Cytosolic
12	LG:027320.5:2002JAN18	914	936		TM	Transmembrane
12	LG:027320.5:2002JAN18	937	1089		TM	Extracellular
12	LG:027320.5:2002JAN18	1	798	-	TM	Extracellular
12	LG:027320.5:2002JAN18	799	821		TM	Transmembrane
12	LG:027320.5:2002JAN18	822	833		TM	Cytosolic
12	LG:027320.5:2002JAN18	834	856		TM	Transmembrane
12	LG:027320.5:2002JAN18	857	865		TM	Extracellular
12	LG:027320.5:2002JAN18	866	888		TM	Transmembrane
12	LG:027320.5:2002JAN18	889	1089		TM	Cytosolic
12	LG:027320.5:2002JAN18	1	185		TM	Cytosolic
12	LG:027320.5:2002JAN18	186	208		TM	Transmembrane
12	LG:027320.5:2002JAN18	209	230		TM	Extracellular
12	LG:027320.5:2002JAN18	231	253		TM	Transmembrane
12	LG:027320.5:2002JAN18	254	265		TM	Cytosolic
12	LG:027320.5:2002JAN18	266	288	_	TM	Transmembrane
		289	309	- 	TM	Extracellular
12	LG:027320.5:2002JAN18	1209	1904		[114]	ILANGONGIGI

TABLE 4

		Total L	lot	IF-2000	Domain	Topology
ł ,	Template ID	Start	Stop	Frame		Topology
NO:		1000	200		Туре	Transmembrane
12	LG:027320.5:2002JAN18	310	332	 	TM	Cytosolic
12	LG:027320.5:2002JAN18	333	352		TM	Transmembrane
12	LG:027320.5:2002JAN18	353	375		TM	Extracellular
12	LG:027320.5:2002JAN18	376	378		TM	
12	LG:027320.5:2002JAN18	379	401		TM	Transmembrane
12	LG:027320.5:2002JAN18	402	772		TM	Cytosolic
12	LG:027320.5:2002JAN18	773	795		TM	Transmembrane
12	LG:027320.5:2002JAN18	796	799		TM	Extracellular
12	LG:027320.5:2002JAN18	800	822		TM	Transmembrane
12	LG:027320.5:2002JAN18	823	834		TM	Cytosolic
12	LG:027320.5:2002JAN18	835	857		TM	Transmembrane
12	LG:027320.5:2002JAN18	858	1089		TM	Extracellular
12	LG:027320.5:2002JAN18	2386	2439	forward 1	SP	
12	LG:027320.5:2002JAN18	2386	2445	forward 1	SP	
12	LG:027320.5:2002JAN18	1129	1188	forward 1	SP	
12	LG:027320.5:2002JAN18	1129	1191	forward 1	SP	
12	LG:027320.5:2002JAN18	2534	2602	forward 2	SP	
12	LG:027320.5:2002JAN18	810	869	forward 3	SP	
12	LG:027320.5:2002JAN18	810	869	forward 3	SP	
12	LG:027320.5:2002JAN18	729	779	forward 3	SP	
12	LG:027320.5:2002JAN18	810	878	forward 3	SP	
13	LG:057499.1:2002JAN18	1	772		TM	Extracellular
13	LG:057499.1:2002JAN18	773	792		TM	Transmembrane
	LG:057499.1:2002JAN18	793	804		TM	Cytosolic
13	LG:057499.1:2002JAN18	805	827		TM	Transmembrane
13	LG:057499.1:2002JAN18	828	1209		TM	Extracellular
13	LG:057499.1:2002JAN18	1210	1232		TM	Transmembrane
13		1233	1290		TM	Cytosolic
13	LG:057499.1:2002JAN18	1291	1310	_	TM	Transmembrane
13	LG:057499.1:2002JAN18	1311	1868		TM	Extracellular
13	LG:057499.1:2002JAN18		1891		TM	Transmembrane
13	LG:057499.1:2002JAN18	1869	1937		TM	Cytosolic
13	LG:057499.1:2002JAN18	1892			TM	Transmembrane
13	LG:057499.1:2002JAN18	1938	1960		TM	Extracellular
13	LG:057499.1:2002JAN18	1961	2243			Extracellular
13	LG:057499.1:2002JAN18	1	933		TNA	Transmembrane
13	LG:057499.1:2002JAN18	934	956		TM	Cytosolic
13	LG:057499.1:2002JAN18	957	1177		TM	Transmembrane
13	LG:057499.1:2002JAN18	1178	1200		TM	
13	LG:057499.1:2002JAN18	1201	1209		TM	Extracellular
13	LG:057499.1:2002JAN18	1210	1232		TM	Transmembrane
13	LG:057499.1:2002JAN18	1233	1345		TM	Cytosolic
13	LG:057499.1:2002JAN18	1346	1368		TM	Transmembrane
13	LG:057499.1:2002JAN18	1369	1387		MT	Extracellular
13	LG:057499.1:2002JAN18	1388	1410		TM	Transmembrane
13	LG:057499.1:2002JAN18	1411	1429		TM	Cytosolic
13	LG:057499.1:2002JAN18	1430	1452		TM	Transmembrane
13	LG:057499.1:2002JAN18	1453	1937		TM	Extracellular
13	LG:057499.1:2002JAN18	1938	1960		TM	Transmembrane
13	LG:057499.1:2002JAN18	1961	2098		TM	Cytosolic

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TABLE 4

		TA	BLE 4				
		Start	Stop	Frame	Do	main	Topology
SEQ ID	Template ID	Sidii	10.05		Typ	ре	
NO:		2099	2121		TM		Transmembrane
13	LG:057499.1:2002JAN18	2122	2140		TIV		Extracellular
13	LG:057499.1:2002JAN18		2163		TM		Transmembrane
13	LG:057499.1:2002JAN18	2141	2169		TN		Cytosolic
13	LG:057499.1:2002JAN18	2164			TIV		Transmembrane
13	LG:057499.1:2002JAN18	2170	2192		TN		Extracellular
13	LG:057499.1:2002JAN18	2193	2243		TN		Extracellular
13	LG:057499.1:2002JAN18	1	771		TN		Transmembrane
13	LG:057499.1:2002JAN18	772	793		TN		Cytosolic
13	LG:057499.1:2002JAN18	794	958		TN		Transmembrane
13	LG:057499.1:2002JAN18	959	981		TN		Extracellular
13	LG:057499.1:2002JAN18	982	1084				Transmembrane
13	LG:057499.1:2002JAN18	1085	1107		T		Cytosolic
13	LG:057499.1:2002JAN18	1108	1200			<u> </u>	Transmembrane
13	LG:057499.1:2002JAN18	1201	1223			<u> </u>	Extracellular
	LG:057499.1:2002JAN18	1224	1303			<u>M</u>	Transmembrane
13	LG:057499.1:2002JAN18	1304	1326			<u>M</u>	
13	LG:057499.1:2002JAN18	1327	1422			<u>M</u>	Cytosolic Transmembrane
13	LG:057499.1:2002JAN18	1423	1445			<u>M</u>	
13	LG:057499.1:2002JAN18	1446	1482			M	Extracellular
13	LG:057499.1:2002JAN18	1483	1505	5		<u>M</u>	Transmembrane
13	LG:05/499.1:20023AN18	1506				M	Cytosolic
13	LG:057499.1:2002JAN18	1594				ſΜ	Transmembrane
13	LG:057499.1:2002JAN18	1614			-	ſΜ	Extracellular
13	LG:057499.1:2002JAN18	1628				TM	Transmembrane
13	LG:057499.1:2002JAN18	1651				MT	Cytosolic
13	LG:057499.1:2002JAN18	1657				TM	Transmembrane
13	LG:057499.1:2002JAN18	1680				TM	Extracellular
13	LG:057499.1:2002JAN18	1684				TM	Transmembrane
13	LG:057499.1:2002JAN18					TM	Cytosolic
13	LG:057499.1:2002JAN18	1707				TM	Transmembrane
13	LG:057499.1:2002JAN18	1800				TM	Extracellular
13	LG:057499.1:2002JAN18	182				TM	Transmembrane
13	LG:057499.1:2002JAN18	193				TM	Cytosolic
13	LG:057499.1:2002JAN18	196				TM	Transmembrane
13	LG:057499.1:2002JAN18	209				TM	Extracellular
13	LG:057499.1:2002JAN18	212				TM	Transmembrane
13	LG:057499.1:2002JAN18	215				TM	Cytosolic
13	LG:057499.1:2002JAN18	217		4Z	rward 1	SP	
13	LG:057499.1:2002JAN18	578				SP	
13	LG:057499.1:2002JAN18	232			rward 1	SP	
13	LG:057499.1:2002JAN18	232			rward 1	SP	
13	LG:057499.1:2002JAN18	232			rward 1	SP SP	
1:	1.G:057499.1:2002JAN18	580			rward 1		
1:	3 IG:057499.1:2002JAN18	589			rward 1	SP	
1		63			rward 3		
1		62			orward 3		
	3 LG:057499.1:2002JAN18	63			orward 3		
1		62	94 6		orward 3		
	0000 IANIZO	63	03 6		orward 3		
				371 fc	orward 3	SP	
	3 LG:057499.1:2002JAN18						

TABLE 4

SEQ ID	Tomplete 15		NOLE 4			
NO:	Template ID	Start	Stop	Frame		Topology
14	10.0(5005.01.0000.11.000		_		Туре	
14	LG:065935.21:2002JAN18	1	166		TM	Extracellular
	LG:065935.21:2002JAN18	167	189		TM	Transmembrane
14	LG:065935.21:2002JAN18	190	201		TM	Cytosolic
14	LG:065935.21:2002JAN18	202	224		TM	Transmembrane
14	LG:065935.21:2002JAN18	225	313		TM	Extracellular
15	LG:071860.12:2002JAN18	1	49		TM	Extracellular
15	LG:071860.12:2002JAN18	50	72		TM	Transmembrane
15	LG:071860.12:2002JAN18	73	117		TM	Cytosolic
15	LG:071860.12:2002JAN18	118	140		TM	Transmembrane
15	LG:071860.12:2002JAN18	141	257		TM	Extracellular
15	LG:071860.12:2002JAN18	240	308	forward 3	SP	
15	LG:071860.12:2002JAN18	240	314	forward 3	SP	
15	LG:071860.12:2002JAN18	240	314	forward 3	SP	
15	LG:071860.12:2002JAN18	240	317	forward 3	SP	
15	LG:071860.12:2002JAN18	240	320	forward 3	SP	
16	LG:087383.29:2002JAN18	1103	1165	forward 2	SP	
16	LG:087383.29:2002JAN18	1100	1153	forward 2	SP	
16	LG:087383.29:2002JAN18	1103	1165	forward 2	SP	
17	LG:098580.3:2002JAN18	1	161		TM	Extracellular
	LG:098580.3:2002JAN18	162	184		TM	Transmembrane
	LG:098580.3:2002JAN18	185	221		TM	Cytosolic
17	LG:098580.3:2002JAN18	222	244		TM	Transmembrane
	LG:098580.3:2002JAN18	245	250		TM	Extracellular
	LG:098580.3:2002JAN18	1	168		TM	Cytosolic
	LG:098580.3:2002JAN18	169	191		TM	Transmembrane
	LG:098580.3:2002JAN18	192	222		TM	Extracellular
	LG:098580.3:2002JAN18	223	245		TM	Transmembrane
	LG:098580.3:2002JAN18	246	250		TM	Cytosolic
	LG:098580.3:2002JAN18	1	219	_	TM	Cytosolic
	LG:098580.3:2002JAN18	220	242			Transmembrane
	LG:098580.3:2002JAN18	243	250		TM	Extracellular
	LG:098580.3:2002JAN18	504	569		SP	
	LG:1001879.1:2002JAN18	148	225		SP	
	LG:1001879.1:2002JAN18	148	225		SP	
	LG:1001879.1:2002JAN18	148	225		SP	
	LG:1001879.1:2002JAN18	148	219		SP	
	LG:1001879.1:2002JAN18	148	207		SP	
	LG:1001879.1:2002JAN18	148	219		SP	
	LG:1001879.1:2002JAN18	148	225		SP	
	LG:1001879.1:2002JAN18	148	210		SP	
	G:1001879.1:2002JAN18	148	204		SP	
	G:1079456.4:2002JAN18	010	218			xtracellular
	G:1079456.4:2002JAN18	219	241			ransmembrane
	G:1079456.4:2002JAN18	242	256			Cytosolic
	G:1079456.4:2002JAN18	548	625		SP	
	G:1079456.4:2002JAN18	548	625		SP	
	.G:1079456.4:2002JAN18 .G:1079456.4:2002JAN18	548	619		SP	
	.G:1079456.4:2002JAN18 .G:1079456.4:2002JAN18	548	601		SP	
17	-C.10/7400.4.2002JAN18	548	625	forward 2	SP _	

TABLE 4

		٦	ABLE	4							
		Start	Į,	top	\[F	rame)	Do	main	Top	ology
SEQ ID	Template ID	Sidii		,iop				Typ	e		
NO:		E 40		513	- 1	forwo		SP			
19	11.6:10/9400.4.20020/ (1110	548				forwo		SP			
19	LG:1079456.4:2002JAN18	548		<u> </u>		forwo		SP			
19	LG:1079456.4:2002JAN18	548		510				SP			
19	LG:1079456.4:2002JAN18	548		619		forwo					
20	LG:1080598.9:2002JAN18	182		238		forwo	ira Z	SP		Evet	racellular
21	LG:1090358.10:2002JAN18	1		3				TIV		Tag	nsmembrane
21	LG:1090358.10:2002JAN18	4		26				TM			
	LG:1090358.10:2002JAN18	27		30				TN			tosolic
21	LG:1090358.10:2002JAN18	31		53				TN			insmembrane
21	LG:1090358.10:2002JAN18	54		733				TN		Ext	racellular
21	LG: 1090358.10:20025A1118	1		3				TN	1		tracellular
21	LG:1090358.10:2002JAN18	4		26				TN	Λ		ansmembrane
21	LG:1090358.10:2002JAN18	27		32				TN	Λ		ytosolic
21	LG:1090358.10:2002JAN18	33		55				TN	1		ansmembrane
21	LG:1090358.10:2002JAN18			733		 		TI		Ex	tracellular
21	LG:1090358.10:2002JAN18	56		128		fona	ard 2			1,	
21	LG:1090358.10:2002JAN18	122					ard 2				
21	LG:1090358.10:2002JAN18	122		127			rard 2	_		_	
21	LG:1090358.10:2002JAN18	122		128		IOIV	uid 2				
21	LG:1090358.10:2002JAN18	122		129			ard 2				
21	LG:1090358.10:2002JAN18	692		757			vard 2				
22	LG:1097492.2:2002JAN18	294	41	301			vard :		P		
22	LG:1097492.2:2002JAN18	29	41	30	12		vard_		P		
	LG:1097492.2:2002JAN18	29	41	30	12		vard	_	P		
22	LG:1097492.2:2002JAN18	29	41	300	06		vard		SP		
22	LG:1097492.2:2002JAN18	53		63	4	for	ward:		SP		
22	LG: 1097492.2.20023A110	23			02	for	ward		SP		
22	LG:1097492.2:2002JAN18		31		.08	for	ward	3	SP		
22	LG:1097492.2:2002JAN18		31		26		ward		SP		
22	LG:1097492.2:2002JAN18		331		102		ward		SP		
22	LG:1097492.2:2002JAN18				396		ward		SP		
22	LG:1097492.2:2002JAN18		331		393		ward		SP		
22	LG:1097492.2:2002JAN18		331_				ward		SP		
22	LG:1097492.2:2002JAN18		331		402	101	waia		TM	-	Extracellular
23	IG:1099945,26:2002JAN18	1			26				TM		Transmembrane
23		7	27		44	}_			TM		Cytosolic
23		3 7	<u>45</u>		42			_	SP		Cytobolic
23		3 1	<u>738 </u>		809		rwarc				
24		1	762		857		rward		SP		
	0.000 1 0.000 1 0.000 1	1	54		58		rward		SP		
24		17	40	8	26		rward		SP		
24			253	1	012		rward		SP		
2			344	1	421		rward		SP		
2			344		421	fc	rwar	2 b	SP		
2			1		374				TM		Extracellular
2	(TO	: 	<u>.</u> 375		397	-			TM		Transmembrane
L	5 LG:1137613.10:2002JAN1		373_ 398		703				TM		Cytosolic
2	5 LG:1137613.10:2002JAN1		70		363 363				TM		Extracellular
2	5 LG:1137613.10:2002JAN1	8	1						TM		Transmembrane
	5 IG:1137613.10:2002JAN1	8	364		386_				TM		Cytosolic
	5 IG:1137613.10:2002JAN1	18	387		398				TM		Transmembrane
	LG:1137613.10:2002JAN	18	399		421				11111		1
<u> </u>											

TABLE 4

		17	ABLE 4			
SEQIE	Template ID	Start	Stop	Frame	Domair	Topology
NO:) Tarrio	Type	Topology
25	LG:1137613.10:2002JAN18	422	903		TM	Extracellular
25	· LG:1137613.10:2002JAN18	1	213		TM	Extracellular
25	LG:1137613.10:2002JAN18	214	236		TM	Transmembrane
25	LG:1137613.10:2002JAN18	237	394		TM	
25	LG:1137613.10:2002JAN18	395	417	- 	TM	Cytosolic
25	LG:1137613.10:2002JAN18	418	458		TM	Transmembrane
25	LG:1137613.10:2002JAN18	459	481			Extracellular
25	LG:1137613.10:2002JAN18	482	665		TM	Transmembrane
25	LG:1137613.10:2002JAN18	666	688		TM	Cytosolic
25	LG:1137613.10:2002JAN18	689	903		TM	Transmembrane
25	LG:1137613.10:2002JAN18	2647	2706	forward 1		Extracellular
25	LG:1137613.10:2002JAN18	2647	2706	forward 1	SP SP	
25	LG:1137613.10:2002JAN18	2647	2703			
25	LG:1137613.10:2002JAN18	2647	2700	forward 1 forward 1	SP	
25	LG:1137613.10:2002JAN18	1028	1087	forward 2	SP	
25	LG:1137613.10:2002JAN18	1172	1258			
25	LG:1137613.10:2002JAN18	1028	1087	forward 2	SP	
25	LG:1137613.10:2002JAN18	1566	1655	forward 2	SP	
25	LG:1137613.10:2002JAN18	1371	1439	forward 3	SP	
25	LG:1137613.10:2002JAN18	1371	1439		SP	
25	LG:1137613.10:2002JAN18	1371	1436	forward 3	SP	
26	LG:118836.26:2002JAN18	3778	3855		SP	
26	LG:118836.26:2002JAN18	1529	1618		SP	
26	LG:118836.26:2002JAN18	1529			SP	
26	LG:118836.26:2002JAN18	1529	1606		SP	
26	LG:118836.26:2002JAN18	4940	1600 5002		SP	
26	LG:118836.26:2002JAN18	1529			SP	
26	LG:118836.26:2002JAN18	3791	1600 3850		SP	
26	LG:118836.26:2002JAN18	756			SP	
26	LG:118836.26:2002JAN18	756	830		SP	-
26	LG:118836.26:2002JAN18	756	809		SP	
26	LG:118836.26:2002JAN18	756	821		SP	
26	LG:118836.26:2002JAN18	+	818		SP	
26	LG:118836.26:2002JAN18	756	815		SP	
27	LG:1330261.32:2002JAN18	756 1	815	forward 3		
27	LG:1330261.32:2002JAN18	 	284			Extracellular
27	LG:1330261.32:2002JAN18	285	307			Transmembrane
27	LG:1330261.32:2002JAN18	308	460			Cytosolic
27	LG:1330261.32:2002JAN18	461	483			Transmembrane
27	LG:1330261.32:2002JAN18	484	511			Extracellular
27	LG:1330261.32:2002JAN18	512	534			ransmembrane .
	LG:1330261.32:2002JAN18	535	559			Cytosolic
	LG:1330261.32:2002JAN18	560	582			ransmembrane (
	LG:1330261.32:2002JAN18	583	616			xtracellular
	LG:1330261.32:2002JAN18	617	639			ransmembrane
	LG:1330261.32:2002JAN18	640	645			Cytosolic
		646	668			ransmembrane
		669	726			xtracellular
	LG:1330261.32:2002JAN18	727	749			ransmembrane
	LG:1330261.32:2002JAN18	750	761	<u> </u>	M (Cytosolic

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		TA	ABLE 4	_			
		Start	Stop	Fro	ame	Domain	Topology
SEQ ID	Template ID	Joran.				Туре	
NO:	911441 2000 20 51 51	762	784			TM	Transmembrane
27	LG:1330261.32:2002JAN18		200			TM	Extracellular
27	LG:1330261.32:2002JAN18		232		rward 1	SP	
27	LG:1330261.32:2002JAN18		927		orward 1	SP	
27	LG:1330261.32:2002JAN18	865	232		orward 1	SP	
27	IG:1330261.32:2002JAN18	2245			orward 1	SP	
27	LG:1330261.32:2002JAN18	4987	504	<u>'</u>	orward 1	SP	
27	IG:1330261.32:2002JAN18	2245	232		orward 1	SP	
27	IG:1330261.32:2002JAN18	3 5104			orward 2	SP	
	IG:1330261.32:2002JAN18	3 1085			orward 2	SP	
	IG:1330261.32:2002JAN18	3 2153			orward 2	SP	
27	LG:1330261.32:2002JAN1	8 [2153				SP	
27	LG:1330261.32:2002JAN1	8 1052			orward 2		
	LG:1330261.32:2002JAN1	8 1052			orward 2		-
27	LG:1330261.32:2002JAN1	8 1052			forward 2		Extracellular
27	LG:1330261.021200 LG:1347461.28:2002JAN1	8 1)51		TM	Transmembrane
28	LG:1347461.28:2002JAN1	8 105	2 10)74		TM	Cytosolic
28	LG:1347461.28:2002JAN	8 107	5 10	085		TM	Transmembrane
28	LG:1347461.28:2002JAN	8 108	6 1	105		TM	
28	LG: 1347401.20.200207 (1			150		TM	Extracellular
28	LG:1347461.28:2002JAN		$\overline{}$	173		TM	Transmembrane
28	LG:1347461.28:2002JAN			282		TM	Cytosolic
28	LG:1347461.28:2002JAN			305		TM	Transmembrane
28	LG:1347461.28:2002JAN			331		TM	Extracellular
28	LG:1347461.28:2002JAN			114	forward	1 SP	
28	LG:1347461.28:2002JAN		•	579		TM	Extracellular
29	LG:1383494.16:2002JAN	118 11		502	 	TM	Transmembrane
29	LG:1383494.16:2002JAN	118 58		712	+	TM	Cytosolic
29	IG:1383494.16:2002JAN	118 100		735	 	TM	Transmembrane
29	IG:1383494.16:2002JAN	118 1/1				TM	Extracellular
29	IG:1383494.16:2002JAN	118 /3		962		TM	Transmembrane
29	IG:1383494.16:2002JA	118 196		985		TM	Cytosolic
29	LG:1383494,16:2002JA	V18 98		1048	ļ	TM	Transmembrane
29		N18 <u> I</u> I)49	1071		TM	Extracellular
	- 1000 404 16:2002 A	N18 11	072	1080			Transmembrane
29		N18 1	081	1103		TM	Cytosolic
20		N18 1	104	1239		TM	Transmembrane
20		N18 1	240	1262		TM	Extracellular
2	10000010	N18 1	263	1276		TM	Transmembrane
2		N18 1	277	1299		TM	
2			300	1441		TM	Cytosolic
		-	442	1464		MT	Transmembran
<u> </u>	9 LG:1383494.16:2002JA		465	1535		TM	Extracellular
	9 LG:1383494.16:2002.J/		1536	1558		TM	Transmembran
2	29 LG:1383494.16:2002J/		1559	1771		TM	Cytosolic
	29 LG:1383494.16:2002J/		1772	1794		TM	Transmembran
1	29 LG:1383494.16:2002J			1800		TM	Extracellular
	20 IG:1383494.16:2002J	AN18	1795	1049		TM	Extracellular
	20 LG:1383494.16:2002J	AN18	1050			TM	Transmembrar
	20 LG:1383494.16:2002J	AN18	1050	1072		TIV	Cytosolic
	20 IG:1383494.16:2002J	AN18	1073	1073		TN	
	29 LG:1383494.16:2002.J	AN18	1074	1096		1110	·

TABLE 4

			TABLE 4			
SEQI	D Template ID	Start	Stop	Frame	10	
NO:		!	Joiop	Inditie	Don	nain Topology
29	LG:1383494.16:2002JAN18	1097	1115		Туре	
29	LG:1383494.16:2002JAN18	1116	1138		TM	Extracellular
29	LG:1383494.16:2002JAN18	1139			TM	Transmembrar
29	LG:1383494.16:2002JAN18		1144		TM	Cytosolic
29	LG:1383494.16:2002JAN18		1164		TM	Transmembran
29	LG:1383494.16:2002JAN18		1253		TM	Extracellular
29	LG:1383494.16:2002JAN18		1276		TM	Transmembran
29	1G:1383404 16:2002 JAN18	1277	1282		TM	Cytosolic
29	LG:1383494.16:2002JAN18		1305		TM	Transmembran
29	LG:1383494.16:2002JAN18	1306	1533		TM	Extracellular
29	LG:1383494.16:2002JAN18	1534	1556		TM	Transmembran
29	LG:1383494.16:2002JAN18	1557	1800		TM	Cytosolic
	LG:1383494.16:2002JAN18	1	263		TM	
29	LG:1383494.16:2002JAN18	264	286		TM	Cytosolic
29	LG:1383494.16:2002JAN18	287	690		TM	Transmembrane
29	LG:1383494.16:2002JAN18	691	709		TM	Extracellular
29	LG:1383494.16:2002JAN18	710	721			Transmembrane
29	LG:1383494.16:2002JAN18	722	744		TM	Cytosolic
29	LG:1383494.16:2002JAN18	745	785		TM	Transmembrane
29	LG:1383494.16:2002JAN18	786	808		TM	Extracellular
29	LG:1383494.16:2002JAN18	809			TM	Transmembrane
29	LG:1383494.16:2002JAN18	958	957		MT	Cytosolic
29	LG:1383494.16:2002JAN18		980		TM	Transmembrane
29	LG:1383494.16:2002JAN18	981	983		TM	Extracellular
29	LG:1383494.16:2002JAN18	984	1003		TM	Transmembrane
29	C:1383404 16:2002 LANDS	1004	1046		TM	Cytosolic
29	LG:1383494.16:2002JAN18	1047	1069		TM	Transmembrane
29	LG:1383494.16:2002JAN18	1070	1078		TM	Extracellular
29	LG:1383494.16:2002JAN18	1079	1101		TM	Transmembrane
29	LG:1383494.16:2002JAN18	1102	1121		TM	Cytosolic
	LG:1383494.16:2002JAN18	1122	1144		TM	Transmembrane
29	LG:1383494.16:2002JAN18	1145	1799		TM	Extracellul
29	LG:1383494.16:2002JAN18	139	225	forward 1	SP	Extracellular
29	LG:1383494.16:2002JAN18	139	225	forward 1	SP	
29	LG:1383494.16:2002JAN18	139	219	forward 1	SP	
29	LG:1383494.16:2002JAN18	139	210			
29	LG:1383494.16:2002JAN18	139	213	forward 1		<u> </u>
29	LG:1383494.16:2002JAN18	139	207	forward 1	SP	
29	LG:1383494.16:2002JAN18	139	198	forward 1	SP	
29	LG:1383494.16:2002JAN18	145	207	forward 1	SP	
29	LG:1383494.16:2002JAN18	4595		forward 1	SP	
29	LG:1383494.16:2002JAN18		4648		SP	
	G:1383494.16:2002JAN18	4595	4654		SP	
	-G:1383494.16:2002JAN18	4595	4651		SP	
	G:1383404 14:2000 (44)16	1692	1751		SP	
	G:1383494.16:2002JAN18	1692	1754	forward 3	SP	
	G:1400155.1:2002JAN18	1	1563		TM	Extracellular
	C.14003 CE 3 0000	1564	1586		TM	Transmembrane
	G:1400155.1:2002JAN18	1587	1644		TM	Cytosolic
30 L	G:1400155.1:2002JAN18]	1041		TM	Extracellular
30 L	G:1400155.1:2002JAN18	1042	1064		TM	Transmembrane
-4/1 II	C.14001EE 2 2222	1065		1	IIVI	Huuisinembrana L

TABLE 4

SEQ ID	Template ID	Start	Stop	Frame		Topology
NO:					Туре	
30	LG:1400155.1:2002JAN18	1202	1224		TM	Transmembrane
30	LG:1400155.1:2002JAN18	1225	1489		TM	Extracellular
30	LG:1400155.1:2002JAN18	1490	1512		TM	Transmembrane
30	LG:1400155.1:2002JAN18	1513	1558		TM	Cytosolic
30	LG:1400155.1:2002JAN18	1559	1581		TM	Transmembrane
30	LG:1400155.1:2002JAN18	1582	1595		TM	Extracellular
30	LG:1400155.1:2002JAN18	1596	1618		TM	Transmembrane
30	LG:1400155.1:2002JAN18	1619	1644		TM	Cytosolic
30	LG:1400155.1:2002JAN18	1	1475		TM	Extracellular
30	LG:1400155.1:2002JAN18	1476	1498		TM	Transmembrane
30	LG:1400155.1:2002JAN18	1499	1552		TM	Cytosolic
30	LG:1400155.1:2002JAN18	1553	1575		TM	Transmembrane
30	LG:1400155.1:2002JAN18	1576	1604		TM	Extracellular
30	LG:1400155.1:2002JAN18	1605	1624		TM	Transmembrane
30	LG:1400155.1:2002JAN18	1625	1643		TM	Cytosolic
30	LG:1400155.1:2002JAN18	2168	2245	forward 2	SP	
30	LG:1400155.1:2002JAN18	2168	2245	forward 2	SP	
30	LG:1400155.1:2002JAN18	2339	2443	forward 2	SP	
30	LG:1400155.1:2002JAN18	2168	2242	forward 2	SP	
30	LG:1400155.1:2002JAN18	2187	2255	forward 3	SP	
30	LG:1400155.1:2002JAN18	2187	2249	forward 3	SP	
30	LG:1400155.1:2002JAN18	2187	2261	forward 3	SP	
30	LG:1400155.1:2002JAN18	2187	2267	forward 3	SP	
30	LG:1400155.1:2002JAN18	2187	2240	forward 3	SP	
31	LG:1446621.1:2002JAN18	1	19		TM	Cytosolic
31	LG:1446621.1:2002JAN18	20	42		TM	Transmembrane
31	LG:1446621.1:2002JAN18	43	353		TM	Extracellular
32	LG:144920.1:2002JAN18	1376	1432	forward 2	SP	
32	LG:144920.1:2002JAN18	1376	1438	forward 2	SP	
32	LG:144920.1:2002JAN18	1376	1435	forward 2	SP	
33	LG:1452619.1:2002JAN18	1	6		TM	Cytosolic
33	LG:1452619.1:2002JAN18	7	29		TM	Transmembrane
33	LG:1452619.1:2002JAN18	30	591		TM	Extracellular
33	LG:1452619.1:2002JAN18	592	614		TM	Transmembrane
33	LG:1452619.1:2002JAN18	615	806		TM	Cytosolic
33	LG:1452619.1:2002JAN18	807	829		TM	Transmembrane
33	LG:1452619.1:2002JAN18	830	838		TM	Extracellular
33	LG:1452619.1:2002JAN18	839	861		TM	Transmembrane
33	LG:1452619.1:2002JAN18	862	872		TM	Cytosolic
33	LG:1452619.1:2002JAN18	873	895		TM	Transmembrane
33	LG:1452619.1:2002JAN18	896	1275		TM	Extracellular
33	LG:1452619.1:2002JAN18	1276	1295		TM	Transmembrane
33	LG:1452619.1:2002JAN18	1296	1323		TM	Cytosolic
33	LG:1452619.1:2002JAN18	1324	1346		TM	Transmembrane
33	LG:1452619.1:2002JAN18	1347	1392		TM	Extracellular
		1393	1412		TM	Transmembrane
33	LG:1452619.1:2002JAN18	1413	1588		TM	Cytosolic
33	LG:1452619.1:2002JAN18	1413	589		TM	Extracellular
33	LG:1452619.1:2002JAN18	500			TM	Transmembrane
33	LG:1452619.1:2002JAN18	590	612		IIVI	Tudibilie

TABLE 4

IABLE 4								
SEQ ID	Template ID	Start	Stop	1	Domain Type	Topology		
NO:	1.0.3.450430.1.2002.IANI38	613	803		TM	Cytosolic		
33	LG:1452619.1:2002JAN18 LG:1452619.1:2002JAN18	804	826		TM	Transmembrane		
33	LG: 1452019.1:2002JAN18	827	835		TM	Extracellular		
33	LG:1452619.1:2002JAN18	836	858		TM	Transmembrane		
33	LG:1452619.1:2002JAN18	859	1329		TM	Cytosolic		
33	LG:1452619.1:2002JAN18	1330	1352		TM	Transmembrane		
33	LG:1452619.1:2002JAN18	1353	1388	 	TM	Extracellular		
33	LG:1452619.1:2002JAN18	1389	1411	 	TM	Transmembrane		
33	LG:1452619.1:2002JAN18	1412	1417		TM	Cytosolic		
33	LG:1452619.1:2002JAN18		1437		TM	Transmembrane		
33	LG:1452619.1:2002JAN18	1418	1506		TM	Extracellular		
33	LG:1452619.1:2002JAN18	1438	1526		TM	Transmembrane		
33	LG:1452619.1:2002JAN18	1507			TM	Cytosolic		
33	LG:1452619.1:2002JAN18	1527	1587		TM	Extracellular		
33	LG:1452619.1:2002JAN18	1	588		TM	Transmembrane		
33	LG:1452619.1:2002JAN18	589	611		TM	Cytosolic		
33	LG:1452619.1:2002JAN18	612	647		TM	Transmembrane		
33	LG:1452619.1:2002JAN18	648	670		TM	Extracellular		
33	LG:1452619.1:2002JAN18	671	679		TM	Transmembrane		
33	LG:1452619.1:2002JAN18	680	697		TM	Cytosolic		
33	LG:1452619.1:2002JAN18	698	759			Transmembrane		
33	LG:1452619.1:2002JAN18	760	782		TM	Extracellular		
33	LG:1452619.1:2002JAN18	783	801		TM	Transmembrane		
33	LG:1452619.1:2002JAN18	802	824		TM	Cytosolic		
33	LG:1452619.1:2002JAN18	825	1270		TM	Transmembrane		
33	LG:1452619.1:2002JAN18	1271	1293		TM	Extracellular		
33	LG:1452619.1:2002JAN18	1294	1296		TM	Transmembrane		
33	LG:1452619.1:2002JAN18	1297	1314		TM			
33	LG:1452619.1:2002JAN18	1315	1330		TM	Cytosolic		
33	LG:1452619.1:2002JAN18	1331	1353		TM	Transmembrane		
33	LG:1452619.1:2002JAN18	1354	1476		TM	Extracellular		
33	LG:1452619.1:2002JAN18	1477	1499		TM	Transmembrane		
33	LG:1452619.1:2002JAN18	1500	1587		TM	Cytosolic		
33	LG:1452619.1:2002JAN18	1780	1851	forward 1				
33	LG:1452619.1:2002JAN18	3805	3888	forward 1				
33	LG:1452619.1:2002JAN18	3043	3099	forward				
33	LG:1452619.1:2002JAN18	902	946	forward 2				
33	LG:1452619.1:2002JAN18	902	952	forward 2				
33	LG:1452619.1:2002JAN18	902	964_	forward 2				
33	LG:1452619.1:2002JAN18	884	955	forward 2				
33	LG:1452619.1:2002JAN18	2111	2176	forward:				
33	LG:1452619.1:2002JAN18	884	982	forward:				
33	LG:1452619.1:2002JAN18	902	982	forward :				
33	LG:1452619.1:2002JAN18	2274	2342	forward				
33	LG:1452619.1:2002JAN18	2253	2342	forward				
34	LG:1453417.6:2002JAN18	1	2453		TM	Extracellular		
34	LG:1453417.6:2002JAN18	2454	2476		TM	Transmembrane		
34	LG:1453417.6:2002JAN18	2477	2488		TM	Cytosolic		
34		2489	2508		TM	Transmembrane		
34		2509	2511		TM	Extracellular		
34	[LG: 1400417.0.20020ANTO	12007	1-20.1					

TABLE 4

050.15	Tanantata ID	Start	Iston	Frame	Domain	Topology
1	Template ID	Sidii	Stop	ridine		Topology
NO:	1 0 1 150 13 7 1 0000 IANI30	0510	0524		Type TM	Transmembrane
	LG:1453417.6:2002JAN18,	2512	2534		TM	Cytosolic
	LG:1453417.6:2002JAN18	2535	2554			Transmembrane
	LG:1453417.6:2002JAN18	2555	2574		TM	Extracellular
	LG:1453417.6:2002JAN18	2575	2593	-	TM	
	LG:1453417.6:2002JAN18	2594	2616		TM	Transmembrane
	LG:1453417.6:2002JAN18	2617	2635		TM	Cytosolic
	LG:1453417.6:2002JAN18	2636	2658		TM	Transmembrane
34	LG:1453417.6:2002JAN18	2659	3378	ļ	TM	Extracellular
34	LG:1453417.6:2002JAN18	3379	3401		TM	Transmembrane
34	LG:1453417.6:2002JAN18	3402	3533		TM	Cytosolic
34	LG:1453417.6:2002JAN18	3534	3556		TM	Transmembrane
34	LG:1453417.6:2002JAN18	3557	3564		TM	Extracellular
34	LG:1453417.6:2002JAN18	1	3534		TM	Extracellular
34	LG:1453417.6:2002JAN18	3535	3557		TM	Transmembrane
34	LG:1453417.6:2002JAN18	3558	3564		TM	Cytosolic
34	LG:1453417.6:2002JAN18	223	324	forward 1	SP	
34	LG:1453417.6:2002JAN18	247	324	forward 1	SP	
34	LG:1453417.6:2002JAN18	259	321	forward 1	SP	
34	LG:1453417.6:2002JAN18	265	318	forward 1	SP	
34	LG:1453417.6:2002JAN18	265	324	forward 1	SP	
34	LG:1453417.6:2002JAN18	7675	7722	forward 1	SP	
34	LG:1453417.6:2002JAN18	5108	5215	forward 2	SP	
34	LG:1453417.6:2002JAN18	4172	4237	forward 2	SP	
34	LG:1453417.6:2002JAN18	7643	7717	forward 2	SP	
34	LG:1453417.6:2002JAN18	7643	7717	forward 2	SP	
34	LG:1453417.6:2002JAN18	5567	5626	forward 2	SP	
34	LG:1453417.6:2002JAN18	7643	7702	forward 2	SP	
34	LG:1453417.6:2002JAN18	1803	1856	forward 3	SP	
35	LG:148485.8:2002JAN18	1	694		TM	Extracellular
35	LG:148485.8:2002JAN18	695	717		TM	Transmembrane
35	LG:148485.8:2002JAN18	718	801		TM	Cytosolic
35	LG:148485.8:2002JAN18	802	824		TM	Transmembrane
35	LG:148485.8:2002JAN18	825	838		TM	Extracellular
35	LG:148485.8:2002JAN18	839	861		TM	Transmembrane
35	LG:148485.8:2002JAN18	862	941		TM	Cytosolic
35	LG:148485.8:2002JAN18	942	964		TM	Transmembrane
35	LG:148485.8:2002JAN18	965	967		TM	Extracellular
35	LG:148485.8:2002JAN18	968	990		TM	Transmembrane
35	LG:148485.8:2002JAN18	991	1069		TM	Cytosolic
35	LG:148485.8:2002JAN18	1070	1089		TM	Transmembrane
35	LG:148485.8:2002JAN18	1090	1114		TM	Extracellular
35	LG:148485.8:2002JAN18	1	20		TM	Cytosolic
35	LG:148485.8:2002JAN18	21	39		TM	Transmembrane
35	LG:148485.8:2002JAN18	40	53		TM	Extracellular
35	LG:148485.8:2002JAN18	54	76		TM	Transmembrane
35	LG:148485.8:2002JAN18	77	192		TM	Cytosolic
35	LG:148485.8:2002JAN18	193	215		TM	Transmembrane
35	LG:148485.8:2002JAN18	216	622		TM	Extracellular
35	LG:148485.8:2002JAN18	623	645		TM	Transmembrane

TABLE 4

C 0 5 15	1		DLE 4	. _[
SEQ ID	Template ID	Start	Stop	Frame	Domain	Topology
NO:					Туре	
35	LG:148485.8:2002JAN18	646	794		TM	Cytosolic
35	LG:148485.8:2002JAN18	795	817		TM	Transmembrane
35	LG:148485.8:2002JAN18	818	845		TM	Extracellular
35	LG:148485.8:2002JAN18	846	868		TM	Transmembrane
35	LG:148485.8:2002JAN18	869	1045		TM	Cytosolic
35	LG:148485.8:2002JAN18	1046	1065		TM	Transmembrane
35	LG:148485.8:2002JAN18	1066	1074		TM	Extracellular
35	LG:148485.8:2002JAN18	1075	1097		TM	Transmembrane
35	LG:148485.8:2002JAN18	1098	1114		TM	Cytosolic
35	LG:148485.8:2002JAN18	1	799		TM	Extracellular
35	LG:148485.8:2002JAN18	800	822		TM	Transmembrane
35	LG:148485.8:2002JAN18	823	834		TM	Cytosolic
35	LG:148485.8:2002JAN18	835	857		TM	Transmembrane
35	LG:148485.8:2002JAN18	858	946		TM	Extracellular
35	LG:148485.8:2002JAN18	947	969		TM	Transmembrane
35	LG:148485.8:2002JAN18	970	980		TM	Cytosolic
35	LG:148485.8:2002JAN18	981	1003		TM	Transmembrane
35	LG:148485.8:2002JAN18	1004	1042	-	TM	Extracellular
35	LG:148485.8:2002JAN18	1043	1065		TM	Transmembrane
35	LG:148485.8:2002JAN18	1066	1113		TM	Cytosolic
35	LG:148485.8:2002JAN18	2381	2452	forward 2	SP	Cylosolic
35	LG:148485.8:2002JAN18	1373	1465	forward 2	SP	
35	LG:148485.8:2002JAN18	1373	1474	forward 2	SP	
35	LG:148485.8:2002JAN18	2381	2446	forward 2	SP	***
36	LG:1502670.1:2002JAN18	293	376	forward 2	SP	
36	LG:1502670.1:2002JAN18	293	382	forward 2	SP	
36	LG:1502670.1:2002JAN18	293	364	forward 2	SP	
36	LG:1502670.1:2002JAN18	293	382	forward 2	SP	
37	LG:206593.3:2002JAN18	1	464	lorward 2	TM	Extracellular
37	LG:206593.3:2002JAN18	465	484		TM	Transmembrane
37	LG:206593.3:2002JAN18	485	516		TM	Cytosolic
37	LG:206593.3:2002JAN18	1364	1462	forward 2	SP	Cytosolic
38	LG:228273.22:2002JAN18	1	924	101Wara 2	TM	Extracellular
38	LG:228273.22:2002JAN18	925	947			Transmembrane
38	LG:228273.22:2002JAN18	948	959		TM	
38	LG:228273.22:2002JAN18	960	982			Cytosolic
38	LG:228273.22:2002JAN18	983	1804		TM TM	Transmembrane
38	LG:228273.22:2002JAN18	1	1643			Extracellular
38	LG:228273.22:2002JAN18	1644			TM	Extracellular
38	LG:228273.22:2002JAN18	1667	1666 1803		TM	Transmembrane
38	· · · · · · · · · · · · · · · · · · ·			for yeard 7	TM	Cytosolic
38	LG:228273.22:2002JAN18	2863	2949	forward 1	SP	
39	LG:228273.22:2002JAN18	3875	3934	forward 2	SP	Fidure - Billion
39	LG:228319.2:2002JAN18	204	305	 	TM	Extracellular
	LG:228319.2:2002JAN18	306	328		TM	Transmembrane
39	LG:228319.2:2002JAN18	329	410		TM	Cytosolic
39	LG:228319.2:2002JAN18	411	433		TM	Transmembrane
39	LG:228319.2:2002JAN18	434	585	I	TM	Extracellular
39	LG:228319.2:2002JAN18	90	152		SP	
39	LG:228319.2:2002JAN18	90	143	forward 3	SP	

TABLE 4

IABLE 4								
SEQ ID	Template ID	Start	Stop	Frame	Domain	Topology		
NO:					Type			
39	LG:228319.2:2002JAN18	90	149	forward 3	SP			
39	LG:228319.2:2002JAN18	90	158	forward 3	SP			
	LG:228319.2:2002JAN18	39	125	forward 3	SP			
39		1	1023	1011111111	TM	Extracellular		
40	LG:229165.16:2002JAN18	1024	1046		TM	Transmembrane		
40	LG:229165.16:2002JAN18		1271		TM	Cytosolic		
40	LG:229165.16:2002JAN18	1047	1294		TM	Transmembrane		
40	LG:229165.16:2002JAN18	1272			TM	Extracellular		
40	LG:229165.16:2002JAN18	1295	1535		TM	Transmembrane		
40	LG:229165.16:2002JAN18	1536	1558					
40	LG:229165.16:2002JAN18	1559	1569		TM	Cytosolic		
40	LG:229165.16:2002JAN18	1570	1587		TM	Transmembrane		
40	LG:229165.16:2002JAN18	1588	1616		TM	Extracellular		
40	LG:229165.16:2002JAN18	1	806		TM	Extracellular		
40	LG:229165.16:2002JAN18	807	826		TM	Transmembrane		
40	LG:229165.16:2002JAN18	827	838		TM	Cytosolic		
40	LG:229165.16:2002JAN18	839	861		TM	Transmembrane		
40	LG:229165.16:2002JAN18	862	1525		TM	Extracellular		
40	LG;229165.16:2002JAN18	1526	1545		TM	Transmembrane		
40	LG:229165.16:2002JAN18	1546	1564		TM	Cytosolic		
40	LG:229165.16:2002JAN18	1565	1587		TM	Transmembrane		
40	LG:229165.16:2002JAN18	1588	1616		TM	Extracellular		
40	LG:229165.16:2002JAN18	1	735		TM	Extracellular ·		
40	LG:229165.16:2002JAN18	736	755		TM	Transmembrane		
40	LG:229165.16:2002JAN18	756	775	- 	TM	Cytosolic		
40	LG:229165.16:2002JAN18	776	798		TM	Transmembrane		
40	LG:229165.16:2002JAN18	799	828		TM	Extracellular		
	LG:229165.16:2002JAN18	829	851	 	TM	Transmembrane		
40		852	862		TM	Cytosolic		
40	LG:229165.16:2002JAN18	863	885		TM	Transmembrane		
40	LG:229165.16:2002JAN18				TM	Extracellular		
40	LG:229165.16:2002JAN18	886	899		TM	Transmembrane		
40	LG:229165.16:2002JAN18	900	931			Cytosolic		
40	LG:229165.16:2002JAN18	932	1270		TM			
40	LG:229165.16:2002JAN18	1271	1293		TM	Transmembrane		
40	LG:229165.16:2002JAN18	1294	1302		TM	Extracellular		
40	LG:229165.16:2002JAN18	1303	1320		TM	Transmembrane		
40	LG:229165.16:2002JAN18	1321	1507		TM	Cytosolic		
40	LG:229165.16:2002JAN18	1508	1530		TM	Transmembrane		
40	LG:229165.16:2002JAN18	1531	1563		TM	Extracellular		
40	LG:229165.16:2002JAN18	1564	1586		TM	Transmembrane		
40	LG:229165.16:2002JAN18	1587	1615		TM	Cytosolic		
40	LG:229165.16:2002JAN18	2080	2166	forward 1				
40	LG:229165.16:2002JAN18	4676	4753	forward 2				
40	LG:229165.16:2002JAN18	4676	4747	forward 2	SP			
40	LG:229165.16:2002JAN18	4676	4753	forward 2	SP			
40	LG:229165.16:2002JAN18	4698	4772	forward 3	SP			
40	LG:229165.16:2002JAN18	4698	4754	forward 3	SP			
40	LG:229165.16:2002JAN18	4698	4757	forward 3				
40	LG:229165.16:2002JAN18	4698	4778	forward 3				
40	LG:229165.16:2002JAN18	4698	4778	forward 3				
40	120,227 100,10,200207 (1410	1.570						

TABLE 4

)LL 4			Tara da anti
SEQ ID	Template ID	Start	Stop	1	ĺ	Topology
NO:			<u> </u>		Туре	
40	LG:229165.16:2002JAN18	1404	1496	1.1.1.1	SP	
41	LG:230895.9:2002JAN18	1	138		TM	Cytosolic
41	LG:230895.9:2002JAN18	139	161		TM	Transmembrane
41	LG:230895.9:2002JAN18	162	596		TM	Extracellular
41	LG:230895.9:2002JAN18	749	820	forward 2	SP	
41	LG:230895.9:2002JAN18	749	826	forward 2	SP	
41	LG:230895.9:2002JAN18	749	820	forward 2	SP	
41	LG:230895.9:2002JAN18	749	814	forward 2	SP	
41	LG:230895.9:2002JAN18	749	826	forward 2	SP	
42	LG:233552.5:2002JAN18	1	1187		TM	Extracellular
42	LG:233552.5:2002JAN18	1188	1210		TM	Transmembrane
42	LG:233552.5:2002JAN18	1211	1228		TM	Cytosolic
42	LG:233552.5:2002JAN18	1	793		TM	Extracellular
42	LG:233552.5:2002JAN18	794	816		TM	Transmembrane
42	LG:233552.5:2002JAN18	817	1132		TM	Cytosolic
42	LG:233552.5:2002JAN18	1133	1152		TM	Transmembrane
42	LG:233552.5:2002JAN18	1153	1161		TM	Extracellular
42	LG:233552.5:2002JAN18	1162	1184	_	TM	Transmembrane
42	LG:233552.5:2002JAN18	1185	1204		TM	Cytosolic
42	LG:233552.5:2002JAN18	1205	1227		TM	Transmembrane
	LG:233552.5:2002JAN18	1228	1228		TM	Extracellular
42	LG:233552.5:2002JAN18	3379	3432	forward 1	SP	
42		3601	3645	forward 1	SP	
42	LG:233552.5:2002JAN18	719	802	forward 2	SP	
42	LG:233552.5:2002JAN18	746	808	forward 2	SP	
42_	LG:233552.5:2002JAN18	746	814	forward 2	SP	
42	LG:233552.5:2002JAN18	719	820	forward 2	SP	
42	LG:233552.5:2002JAN18	170	223	forward 2	SP	
42	LG:233552.5:2002JAN18		229	forward 2	SP	
42	LG:233552.5:2002JAN18	170	2298	forward 1	SP	
43	LG:234430.7:2002JAN18	2233	2292	forward 1	SP	
43	LG:234430.7:2002JAN18	2233			SP	
43	LG:234430.7:2002JAN18	2233	2346	forward 1		
43	LG:234430.7:2002JAN18	1817	1894	forward 2		
43	LG:234430.7:2002JAN18	1817	1894	forward 2		
43	LG:234430.7:2002JAN18	1874	1945	forward 2		Cytosolic
44	LG:236659.1:2002JAN18	1	155		TM	Transmembrane
44	LG:236659.1:2002JAN18	156	178		TM	
44	LG:236659.1:2002JAN18	179	1635		TM	Extracellular
44	LG:236659.1:2002JAN18	1636	1658		TM	Transmembrane
44	LG:236659.1:2002JAN18	1659	1678		TM	Cytosolic
44	LG:236659.1:2002JAN18	1679	1701		TM	Transmembrane
44	LG:236659.1:2002JAN18	1702	1715		TM	Extracellular
44	LG:236659.1:2002JAN18	1716	1734		TM	Transmembrane
44	LG:236659.1:2002JAN18	1735	1740		TM	Cytosolic
44	LG:236659.1:2002JAN18	1741	1760		TM	Transmembrane
44	LG:236659.1:2002JAN18	1761	1794		TM	Extracellular
44	LG:236659.1:2002JAN18	1795	1817		TM_	Transmembrane
44	LG:236659.1:2002JAN18	1818	1873		TM	Cytosolic
44	LG:236659.1:2002JAN18	1874	1893		TM	Transmembrane

TABLE 4

050 10	1-1-15	Start	Cton	Frame	Domain	Topology
SEQ ID	Template ID	Sidii	Stop	ł I	Type	lopology
NO:	1.0.00((50.1.0000 IANII	1894	2001		TM	Extracellular
44	LG:236659.1:2002JAN18	2002	2024		TM	Transmembrane
44	LG:236659.1:2002JAN18	2002	2024		TM	Cytosolic
44	LG:236659.1:2002JAN18	2025	2116		TM	Transmembrane
44	LG:236659.1:2002JAN18	2117	2334		TM	Extracellular
44	LG:236659.1:2002JAN18	2335	2357		TM	Transmembrane
44	LG:236659.1:2002JAN18	2358	2377		TM	Cytosolic
44	LG:236659.1:2002JAN18				TM	Transmembrane
44	LG:236659.1:2002JAN18	2378	2395		TM	Extracellular
44	LG:236659.1:2002JAN18	2396	2440		TM	Transmembrane
44	LG:236659.1:2002JAN18	2441	2459		TM	Cytosolic
44	LG:236659.1:2002JAN18	2460	2495		TM	Transmembrane
44	LG:236659.1:2002JAN18	2496	2518		TM	Extracellular
44	LG:236659.1:2002JAN18	2519	2915			
44	LG:236659.1:2002JAN18	1	1676		TM	Extracellular
44	LG:236659.1:2002JAN18	1677	1696		TM	Transmembrane
44	LG:236659.1:2002JAN18	1697	1708		TM	Cytosolic
44	LG:236659.1:2002JAN18	1709	1731		TM	Transmembrane
44	LG:236659.1:2002JAN18	1732	1740		TM	Extracellular
44	LG:236659.1:2002JAN18	1741	1760		TM	Transmembrane
44	LG:236659.1:2002JAN18	1761	1780		TM	Cytosolic
44	LG:236659.1:2002JAN18	1781	1803		TM	Transmembrane
44	LG:236659.1:2002JAN18	1804	2473		TM	Extracellular
44	LG:236659.1:2002JAN18	2474	2496		TM	Transmembrane
44	LG:236659.1:2002JAN18	2497	2502		TM	Cytosolic
44	LG:236659.1:2002JAN18	2503	2525		TM	Transmembrane
44	LG:236659.1:2002JAN18	2526	2914		TM	Extracellular
44	LG:236659.1:2002JAN18	1	1736		TM	Extracellular
44	LG:236659.1:2002JAN18	1737	1759		TM	Transmembrane
44	LG:236659.1:2002JAN18	1760	1993		TM	Cytosolic
44	LG:236659.1:2002JAN18	1994	2016		TM	Transmembrane
44	LG:236659.1:2002JAN18	2017	2020		TM	Extracellular
44	LG:236659.1:2002JAN18	2021	2038		TM	Transmembrane
44	LG:236659.1:2002JAN18	2039	2335	<u> </u>	TM	Cytosolic
44	LG:236659.1:2002JAN18	2336	2358		TM	Transmembrane
44	LG:236659.1:2002JAN18	2359	2914		TM	Extracellular
44	LG:236659.1:2002JAN18	5200	5262	forward 1	SP	
44	LG:236659.1:2002JAN18	3833	3895		SP	
44	LG:236659.1:2002JAN18	6732	6797		SP	
44	LG:236659.1:2002JAN18	1026	1106		SP	
44	LG:236659.1:2002JAN18	1026	1100		SP	
45	LG:236767.26:2002JAN18	1118	1189		SP	
45	LG:236767.26:2002JAN18	1118	1195		SP	
45	LG:236767.26:2002JAN18	1118	1189	forward 2	SP	
45	LG:236767.26:2002JAN18	1118	1183	forward 2	SP	<u></u>
46	LG:237489.7:2002JAN18	1	70		TM	Extracellular
46	LG:237489.7:2002JAN18	71	93		TM	Transmembrane
46	LG:237489.7:2002JAN18	94	94		TM	Cytosolic
46	LG:237489.7:2002JAN18	95	117		TM	Transmembrane
46	LG:237489.7:2002JAN18	118	1144		TM	Extracellular

TABLE 4

SEQ ID	Template ID	Start	Stop	Frame	Domain	Topology
NO:		Jack	SiOp	ridine	Туре	Topology
46	LG:237489.7:2002JAN18	1	1123	- 	TM	Extracellular
46	LG:237489.7:2002JAN18	1124	1143		TM	Transmembrane
	LG:237489.7:2002JAN18	1144	1143		TM	Cytosolic
46		1144	11113		TM	Extracellular
46	LG:237489.7:2002JAN18				TM	
46	LG:237489.7:2002JAN18	1114	1136		TM	Transmembrane
46	LG:237489.7:2002JAN18	1137	1143	(fam. 1000)	SP	Cytosolic
46	LG:237489.7:2002JAN18	1096	1152	forward 1	SP	
46	LG:237489.7:2002JAN18	1568	1684	forward 2		
46	LG:237489.7:2002JAN18	2484	2564	forward 3	SP	
46	LG:237489.7:2002JAN18	2484	2558	forward 3	SP	
46	LG:237489.7:2002JAN18	2484	2567	forward 3	SP	
46	LG:237489.7:2002JAN18	2484	2555	forward 3	SP	
46	LG:237489.7:2002JAN18	2484	2573	forward 3	SP	
46	LG:237489.7:2002JAN18	2484	2549	forward 3	SP	
46	LG:237489.7:2002JAN18	2484	2564	forward 3	SP	
47	LG:238218.20:2002JAN18	1	794		TM	Extracellular
47	LG:238218.20:2002JAN18	795	817		TM	Transmembrane
47	LG:238218.20:2002JAN18	818	1053		TM	Cytosolic
47	LG:238218.20:2002JAN18	1054	1076		TM	Transmembrane
47	LG:238218.20:2002JAN18	1077	1090		TM	Extracellular
47	LG:238218.20:2002JAN18	1091	1113		TM	Transmembrane
47	LG:238218.20:2002JAN18	1114	1265		TM	Cytosolic
47	LG:238218.20:2002JAN18	1266	1288		TM	Transmembrane
47	LG:238218.20:2002JAN18	1289	1630		TM	Extracellular
47	LG:238218.20:2002JAN18	1631	1653		TM	Transmembrane
47	LG:238218.20:2002JAN18	1654	1735		TM	Cytosolic
47	LG:238218.20:2002JAN18	1736	1758		TM	Transmembrane
47	LG:238218.20:2002JAN18	1759	1761		TM	Extracellular
47	LG:238218.20:2002JAN18	1762	1779		TM	Transmembrane
47	LG:238218.20:2002JAN18	1780	1799		TM	Cytosolic
47	LG:238218.20:2002JAN18	1800	1822		TM	Transmembrane
47	LG:238218.20:2002JAN18	1823	1836		TM	Extracellular
47	LG:238218.20:2002JAN18	1837	1859		TM	Transmembrane
47	LG:238218.20:2002JAN18	1860	2005		TM	Cytosolic
47	LG:238218.20:2002JAN18	1	1584		TM	Extracellular
47	LG:238218.20:2002JAN18	1585	1607		TM	Transmembrane
47	LG:238218.20:2002JAN18	1608	1639		TM	Cytosolic
47	LG:238218.20:2002JAN18	1640	1662		TM	Transmembrane
47	LG:238218.20:2002JAN18	1663	2005		TM	Extracellular
47	LG:238218.20:2002JAN18	1	1090		TM	Extracellular
47	LG:238218.20:2002JAN18	1091	1113		TM	Transmembrane
47	LG:238218.20:2002JAN18	1114	1142		TM	Cytosolic
47	LG:238218.20:2002JAN18	1143	1165		TM	Transmembrane
47	LG:238218.20:2002JAN18	1166	1222		TM	Extracellular
47	LG:238218.20:2002JAN18	1223	1245		TM	Transmembrane
47	LG:238218.20:2002JAN18	1246	1265		TM	Cytosolic
		1266	1288		TM	Transmembrane
47	LG:238218.20:2002JAN18		1793		TM	Extracellular
47	LG:238218.20:2002JAN18	1289				
47	LG:238218.20:2002JAN18	1794	1816		TM	Transmembrane

TABLE 4

			DLE 4			
SEQ ID NO:	Template ID	Start	Stop	Frame	Domain Type	Topology
47	LG:238218.20:2002JAN18	1817	1836		TM	Cytosolic
47	LG:238218.20:2002JAN18	1837	1859		TM	Transmembrane
47	LG:238218.20:2002JAN18	1860	2005		TM	Extracellular
47	LG:238218.20:2002JAN18	3814	3900	forward 1	SP	
47	LG:238218.20:2002JAN18	3814	3879	forward 1	SP	
47	LG:238218.20:2002JAN18	3814	3891	forward 1	SP	
47	LG:238218.20:2002JAN18	4927	4980	forward 1	SP	
47	LG:238218.20:2002JAN18	3814	3891	forward 1	SP	
47	LG:238218.20:2002JAN18	4927	4974	forward 1	SP	
47	LG:238218.20:2002JAN18	3814	3885	forward 1	SP	
47	LG:238218.20:2002JAN18	4927	4989	forward 1	SP	
47	LG:238218.20:2002JAN18	4900	4992	forward 1	SP	
47	LG:238218.20:2002JAN18	4927	4980	forward 1	SP	
47	LG:238218.20:2002JAN18	3814	3885	forward 1	SP	
47	LG:238218.20:2002JAN18	518	598	forward 2	SP	
47	LG:238218.20:2002JAN18	518	598	forward 2	SP	
47	LG:238218.20:2002JAN18	518	583	forward 2	SP	
47	LG:238218.20:2002JAN18	518	586	forward 2	SP	
47	LG:238218.20:2002JAN18	518	601	forward 2	SP	
47	LG:238218.20:2002JAN18	518	565	forward 2	SP	
47	LG:238218.20:2002JAN18	518	571	forward 2	SP	
47	LG:238218.20:2002JAN18	2400	2456	forward 3	SP	
47	LG:238218.20:2002JAN18	2400	2459	forward 3	SP	
47	LG:238218.20:2002JAN18	1941	2009	forward 3	SP	
47	LG:238218.20:2002JAN18	1941	2015	forward 3	SP	
47	LG:238218.20:2002JAN18	1941	2003	forward 3	SP	
47	LG:238218.20:2002JAN18	1941	2015	forward 3	SP	
47	LG:238218.20:2002JAN18	1941	2015	forward 3	SP	
48	LG:239939.14:2002JAN18	1	3		TM	Extracellular.
48	LG:239939.14:2002JAN18	4	26		TM	Transmembrane
48	LG:239939.14:2002JAN18	27	149		TM	Cytosolic
48	LG:239939.14:2002JAN18	150	169		TM	Transmembrane
48	LG:239939.14:2002JAN18	170	183		TM	Extracellular
48	LG:239939.14:2002JAN18	184	206		TM	Transmembrane
48	LG:239939.14:2002JAN18	207	379		TM	Cytosolic
48	LG:239939.14:2002JAN18	380	402	_	TM	Transmembrane
48	LG:239939.14:2002JAN18	403	434		TM	Extracellular
48	LG:239939.14:2002JAN18	873	926	forward 3		
48	LG:239939.14:2002JAN18	873	929	forward 3		
48	LG:239939.14:2002JAN18	873	956	forward 3		
49	LG:242288.11:2002JAN18	973	1044	forward 1		
49	LG:242288.11:2002JAN18	4097	4195	forward 2		
50	LG:242491.29:2002JAN18	64	117	forward 1		
50	LG:242491.29:2002JAN18	64	120	forward 1		
50	LG:242491.29:2002JAN18	64	123	forward 1		
50	LG:242491.29:2002JAN18	31	117	forward 1		
50	LG:242491.29:2002JAN18	64	126	forward 1		
50	LG:242491.29:2002JAN18	64	117	forward 1		
	LG:242491.29:2002JAN18	31	123	forward 1		
50	LG:242491.29:2002JAN18	31	1123	polwara 1	Jor	

TABLE 4

	Template ID	Start	Stop	Frame	Domain	Topology
NO:					Туре	
50	LG:242491.29:2002JAN18	64	129	forward 1	SP	
50	LG:242491.29:2002JAN18	43	117	forward 1	SP	
50	LG:242491.29:2002JAN18	31	123	forward 1	SP	
51	LG:243488.41:2002JAN18	1	525		TM	Extracellular
51	LG:243488.41:2002JAN18	526	548		TM	Transmembrane
51	LG:243488.41:2002JAN18	549	655		TM	Cytosolic
51	LG:243488.41:2002JAN18	656	678		TM	Transmembrane
51	LG:243488.41:2002JAN18	679	715		TM	Extracellular
51	LG:243488.41:2002JAN18	716	735		TM	Transmembrane
51	LG:243488.41:2002JAN18	736	742		TM	Cytosolic
51	LG:243488.41:2002JAN18	1	665		TM	Extracellular
51	LG:243488.41:2002JAN18	666	688		TM	Transmembrane
51	LG:243488.41:2002JAN18	689	742		TM	Cytosolic
51	LG:243488.41:2002JAN18	361	438	forward 1	SP	
51	LG:243488.41:2002JAN18	1940	2017	forward 2	SP	
51	LG:243488.41:2002JAN18	1940	2020	forward 2	SP	
52	LG:247792.18:2002JAN18	1	1050		TM	Extracellular
52	LG:247792.18:2002JAN18	1051	1073		TM	Transmembrane
52	LG:247792.18:2002JAN18	1074	1142	-	TM	Cytosolic
52	LG:247792.18:2002JAN18	1930	2016	forward 1	SP	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,
52	LG:247792.18:2002JAN18	1930	2010	forward 1	SP	· · · · · · · · · · · · · · · · · · ·
53	LG:253193.17:2002JAN18	1	201		TM	Extracellular
53	LG:253193.17:2002JAN18	202	224		TM	Transmembrane
53	LG:253193.17:2002JAN18	225	440		TM	Cytosolic
53	LG:253193.17:2002JAN18	441	463		TM	Transmembrane
53	LG:253193.17:2002JAN18	464	771	· · · · · · · · · · · · · · · · · · ·	TM	Extracellular
53	LG:253193.17:2002JAN18	11	52		TM	Cytosolic
53	LG:253193.17:2002JAN18	53	75		TM	Transmembrane
53	LG:253193.17:2002JAN18	76	84		TM	Extracellular
53	LG:253193.17:2002JAN18	85	107		TM	Transmembrane
53	LG:253193.17:2002JAN18	108	200		TM	Cytosolic
53	LG:253193.17:2002JAN18	201	223		TM	Transmembrane
53	LG:253193.17:2002JAN18	224	291		TM	Extracellular
53	LG:253193.17:2002JAN18	292	314		TM	Transmembrane
53	LG:253193.17:2002JAN18	315	371		TM	Cytosolic
53	LG:253193,17:2002JAN18	372	394		TM	Transmembrane
53	LG:253193.17:2002JAN18	395	770		TM	Extracellular
53	LG:253193.17:2002JAN18	1	20		TM	Cytosolic
53	LG:253193.17:2002JAN18	21	40		TM	Transmembrane
53	LG:253193.17:2002JAN18	41	49	 	TM	Extracellular
53	LG:253193.17:2002JAN18	50	72		TM	Transmembrane
53	LG:253193.17:2002JAN18	73	152		TM	Cytosolic
53	LG:253193.17:2002JAN18	153	171		TM	Transmembrane
53	LG:253193.17:2002JAN18	172	211		TM	Extracellular
53	LG:253193.17:2002JAN18	212	234		TM	Transmembrane
53	LG:253193.17:2002JAN18	235	290		TM	Cytosolic
53	LG:253193.17:2002JAN18	291	313		TM	Transmembrane
53	LG:253193.17:2002JAN18	314	770		TM	Extracellular
53	LG:253193.17:2002JAN18	1513		fongard 1	SP	LANGCENGIG
	119.200140.17.2002JAIN10	11010	1578	forward 1	Jor	<u> </u>

TABLE 4

TABLE 4								
SEQ ID T	remplate ID	Start	Stop	Frame		Topology		
NO:					Туре			
53	(5 .200170,17.20020,	257	322	forward 2	SP			
53	LG:253193.17:2002JAN18	257	319	forward 2	SP	Cutopolic		
54	LG:257088.20:2002JAN18	1	20		TM	Cytosolic		
54	LG:257088.20:2002JAN18	21	43		TM	Transmembrane		
54	LG:257088.20:2002JAN18	44	1253		TM	Extracellular		
	LG:257088.20:2002JAN18	1028	1153	forward 2	SP			
54	LG:257088.20:2002JAN18	1028	1147	forward 2	SP			
	LG:257088.20;2002JAN18	1937	1999	forward 2	SP			
	LG:265552.1:2002JAN18	ī	218		TM	Cytosolic		
	LG:265552.1:2002JAN18	219	238		TM	Transmembrane		
55	LG:265552.1:2002JAN18	239	247		TM	Extracellular		
55	LG:265552.1:2002JAN18	248	267		TM	Transmembrane		
55	LG:265552.1:2002JAN18	268	341		TM	Cytosolic		
55	LG:265552.1:2002JAN18	342	364		TM	Transmembrane		
55	LG:265552.1:2002JAN18	365	630		TM	Extracellular		
55	LG:265552.1:2002JAN18	1	163		TM	Extracellular		
55	LG:265552.1:2002JAN18	164	186		TM	Transmembrane		
55	LG:265552.1:2002JAN18	187	226		TM	Cytosolic		
55	LG:205552.1:2002JAN18	227	249		TM	Transmembrane		
55	LG:265552.1:2002JAN18	250	291		TM	Extracellular		
55	LG:265552.1:2002JAN18	292	314		TM	Transmembrane		
55	LG:265552.1:2002JAN18	315	629		TM	Cytosolic		
55	LG:265552.1:2002JAN18	1	381		TM	Extracellular		
55	LG:265552.1:2002JAN18	382	404		TM	Transmembrane		
55	LG:265552.1:2002JAN18	405	483		TM	Cytosolic		
55	LG:265552.1:2002JAN18	484	506		TM	Transmembrane		
55	LG:265552.1:2002JAN18	507	629		TM	Extracellular		
55	LG:265552.1:2002JAN18	817	867	forward				
55	LG:265552.1:2002JAN18	683	754	forward				
55	LG:265552.1:2002JAN18		742	forward				
. 55	LG:265552.1:2002JAN18	683	760	forward				
55	LG:265552.1:2002JAN18	683	748	forward				
55	LG:265552.1:2002JAN18	683	757	forward				
55	LG:265552.1:2002JAN18	683	745	forward				
55	LG:265552.1:2002JAN18	683		forward				
55	LG:265552.1:2002JAN18	683	748_	- Ioiwaia	TM	Extracellular		
56	LG:275355.12:2002JAN18	1 007	386		TM	Transmembran		
56	LG:275355.12:2002JAN18	387	409		TM	Cytosolic		
56	LG:275355.12:2002JAN18	410	429		TM	Transmembran		
56	LG:275355.12:2002JAN18	430	452		TM	Extracellular		
56	LG:275355.12:2002JAN18	453	514		TM	Extracellular		
56	LG:275355.12:2002JAN18	1	387		TM	Transmembran		
56	LG:275355.12:2002JAN18	388	410		TM	Cytosolic		
56	LG:275355.12:2002JAN18	411	429		TM	Transmembrar		
56	LG:275355.12:2002JAN18	430	452			Extracellular		
56	LG:275355.12:2002JAN18	453	513		MT	LATIGOGIIGIGI		
56	LG:275355.12:2002JAN18	1084						
56	LG:275355.12:2002JAN18	1084						
56		1084		forward				
	LG:275355.12:2002JAN18	86	145	I for your	コンコベレ	ı		

TABLE 4

SEQ ID	Township ID	lo: ·				
i i	Template ID	Start	Stop	Frame		Topology
NO:	1.0.000014.1.000044.110	_ <u> </u>			Туре	
57	LG:280014.1:2002JAN18	1	100		TM	Extracellular
57	LG:280014.1:2002JAN18	101	123		TM	Transmembrane
57	LG:280014.1:2002JAN18	124	238		TM	Cytosolic
57	LG:280014.1:2002JAN18	239	256		TM	Transmembrane
57	LG:280014.1:2002JAN18	257	270		TM	Extracellular
57	LG:280014.1:2002JAN18	271	293		TM	Transmembrane
57	LG:280014.1:2002JAN18	294	314		TM	Cytosolic
57	LG:280014.1:2002JAN18	315	334		TM	Transmembrane
57	LG:280014.1:2002JAN18	335	335		TM	Extracellular
57	LG:280014.1:2002JAN18	1	99		TM	Extracellular
57	LG:280014.1:2002JAN18	100	122		TM	Transmembrane
57	LG:280014.1:2002JAN18	123	335		TM'	Cytosolic
57	LG:280014.1:2002JAN18	457	543	forward 1	SP	
57	LG:280014.1:2002JAN18	457	537	forward 1	SP	
57	LG:280014.1:2002JAN18	457	528	forward 1	SP	
57	LG:280014.1:2002JAN18	457	528	forward 1	SP	
57	LG:280014.1:2002JAN18	457	543	forward 1	SP	
57	LG:280014.1:2002JAN18	712	765	forward 1	SP	
57	LG:280014.1:2002JAN18	457	549	forward 1	SP	
57	LG:280014.1:2002JAN18	165	239	forward 3	SP	
57	LG:280014.1:2002JAN18	165	236	forward 3	SP	
57	LG:280014.1:2002JAN18	165	236	forward 3	SP	
57	LG:280014.1:2002JAN18	165	251	forward 3	SP	
57	LG:280014.1:2002JAN18	165	242	forward 3	SP	
57	LG:280014.1:2002JAN18	267	335	forward 3	SP	
58	LG:299937.3:2002JAN18	1	790		TM	Extracellular
58	LG:299937.3:2002JAN18	791	813		TM	Transmembrane
58	LG:299937.3:2002JAN18	814	1010		TM	Cytosolic
58	LG:299937.3:2002JAN18	817	888	forward 1	SP	CY1000IIC
58	LG:299937.3:2002JAN18	2726	2785	forward 2	SP	
59	LG:311197.3:2002JAN18	1	1547		TM	Extracellular
59	LG:311197.3:2002JAN18	1548	1570		TM	Transmembrane
59	LG:311197.3:2002JAN18	1571	1584		TM	Cytosolic
59	LG:311197.3:2002JAN18	2227	2307	forward 1	SP	Cy1030IIC
59	LG:311197.3:2002JAN18	2227	2322	forward 1	SP	
59	LG:311197.3:2002JAN18	4552	4608	forward 1	SP	
	LG:311197.3:2002JAN18	4552	4614	forward 1	SP	
59	LG:311197.3:2002JAN18	2850	2915		SP	
59	LG:311197.3:2002JAN18	2850	2921		SP	
59	LG:311197.3:2002JAN18	675	761		SP	
	LG:321069.2:2002JAN18	1	630	70.114.0		Extracellular
	LG:321069.2:2002JAN18	631	653	<u> </u>		Transmembrane
	LG:321069.2:2002JAN18	654	665	 	TM	Cytosolic
	LG:321069.2:2002JAN18	666	688	-		Transmembrane
	LG:321069.2:2002JAN18	689	702	- 		Extracellular
	LG:321069.2:2002JAN18	703	725	- 		
	LG:321069.2:2002JAN18	726	886			Transmembrane
	LG:321069.2:2002JAN18	887	909			Cytosolic
	LG:321069.2:2002JAN18	910	1343			Transmembrane
	20.0210071212002071110	710	1040		TM	Extracellular

TABLE 4

SEQ ID	Template ID	Start	Stop	Frame	Domain	Topology
NO:	Template 12	0,0	0.00		Туре	,
60	LG:321069.2:2002JAN18	1	. 634		TM	Extracellular
60	LG:321069.2:2002JAN18	635	653		TM	Transmembrane
60	LG:321069.2:2002JAN18	654	664		TM	Cytosolic
60	LG:321069.2:2002JAN18	665	687		TM	Transmembrane
60	LG:321069.2:2002JAN18	688	806		TM	Extracellular
60	LG:321069.2:2002JAN18	807	829		TM	Transmembrane
60	LG:321069.2:2002JAN18	830	1078		TM	Cytosolic
60	LG:321069.2:2002JAN18	1079	1101		TM	Transmembrane
60	LG:321069.2:2002JAN18	1102	1127		TM	Extracellular
60	LG:321069.2:2002JAN18	1128	1150		TM	Transmembrane
60	LG:321069.2:2002JAN18	1151	1343		TM	Cytosolic
60	LG:321069.2:2002JAN18	1	448		TM	Cytosolic
60	LG:321069.2:2002JAN18	449	468		TM	Transmembrane
60	LG:321069.2:2002JAN18	469	477		TM	Extracellular
60	LG:321069.2:2002JAN18	478	500		TM	Transmembrane
60	LG:321069.2:2002JAN18	501	506		TM	Cytosolic
60	LG:321069.2:2002JAN18	507	529		TM	Transmembrane
60	LG:321069.2:2002JAN18	530	630		TM	Extracellular
60	LG:321069.2:2002JAN18	631	653		TM	Transmembrane
60	LG:321069.2:2002JAN18	654	665		TM	Cytosolic
60	LG:321069.2:2002JAN18	666	688		TM	Transmembrane
60	LG:321069.2:2002JAN18	689	692		TM	Extracellular
60	LG:321069.2:2002JAN18	693	712		TM	Transmembrane
60	LG:321069.2:2002JAN18	713	1070		TM	Cytosolic
60	LG:321069.2:2002JAN18	1071	1093		TM	Transmembrane
60	LG:321069.2:2002JAN18	1094	1102		TM	Extracellular
60	LG:321069.2:2002JAN18	1103	1122		TM	Transmembrane
60	LG:321069.2:2002JAN18	1123	1128		TM	Cytosolic
60	LG:321069.2:2002JAN18	1129	1151		TM	Transmembrane
60	LG:321069.2:2002JAN18	1152	1343		TM	Extracellular
60	LG:321069.2:2002JAN18	535	600	forward 1	SP	
60	LG:321069.2:2002JAN18	535	606	forward 1	SP	
61	LG:330900.8:2002JAN18	1	1252		TM	Extracellular
61	LG:330900.8:2002JAN18	1253	1275		TM	Transmembrane
61	LG:330900.8:2002JAN18	1276	1294		TM	Cytosolic
61	LG:330900.8:2002JAN18	1295	1317		TM	Transmembrane
61	LG:330900.8:2002JAN18	1318	1326		TM	Extracellular
61	LG:330900.8:2002JAN18	1327	1349		TM	Transmembrane
61	LG:330900.8:2002JAN18	1350	1360		TM	Cytosolic
61	LG:330900.8:2002JAN18	1361	1383		TM	Transmembrane
61	LG:330900.8:2002JAN18	1384	1458		TM	Extracellular
61	LG:330900.8:2002JAN18	1459	1481		TM	Transmembrane
61	LG:330900.8:2002JAN18	1482	1501		TM	Cytosolic
61	LG:330900.8:2002JAN18	1502	1524		TM	Transmembrane
61	LG:330900.8:2002JAN18	1525	1577		TM	Extracellular
61	LG:330900.8:2002JAN18	1578	1600		TM	Transmembrane
61	LG:330900.8:2002JAN18	1601	1630		TM	Cytosolic
61	LG:330900.8:2002JAN18	1631	1653		TM	Transmembrane
61	LG:330900.8:2002JAN18	1654	1678		TM	Extracellular

TABLE 4

		170	SLE 4			
1	Template ID	Start	Stop	1	Domain Type	Topology
NO:	1.0.222222.2.2022.14.113.8	+	1649		TM	Extracellular
61	LG:330900.8:2002JAN18	1650	1672		TM	Transmembrane
61	LG:330900.8:2002JAN18	1673	1678		TM	Cytosolic
61	LG:330900.8:2002JAN18	10/3	22		TM	Extracellular
61	LG:330900.8:2002JAN18	23	45		TM	Transmembrane
61	LG:330900.8:2002JAN18	46	115		TM	Cytosolic
61	LG:330900.8:2002JAN18	116	135		TM	Transmembrane
61	LG:330900.8:2002JAN18		437		TM	Extracellular
61	LG:330900.8:2002JAN18	136			TM	Transmembrane
61	LG:330900.8:2002JAN18	438	460		TM	Cytosolic
61	LG:330900.8:2002JAN18	461	596		TM	Transmembrane
61	LG:330900.8:2002JAN18	597	619		TM	Extracellular
61	LG:330900.8:2002JAN18	620	633		TM	Transmembrane
61	LG:330900.8:2002JAN18	634	651			Cytosolic
61	LG:330900.8:2002JAN18	652	671		TM	Transmembrane
61	LG:330900.8:2002JAN18	672	694		TM	
61	LG:330900.8:2002JAN18	695	1356		TM	Extracellular Transmembrane
61	LG:330900.8:2002JAN18	1357	1376		TM	
61	LG:330900.8:2002JAN18	1377	1380		TM	Cytosolic
61	LG:330900.8:2002JAN18	1381	1403		TM	Transmembrane
61	LG:330900.8:2002JAN18	1404	1433		TM	Extracellular
61	LG:330900.8:2002JAN18	1434	1456		TM	Transmembrane
61	LG:330900.8:2002JAN18	1457	1476		TM	Cytosolic
61	LG:330900.8:2002JAN18	1477	1499		MT	Transmembrane
61	LG:330900.8:2002JAN18	1500	1503		TM	Extracellular
61	LG:330900.8:2002JAN18	1504	1526		TM	Transmembrane
61	LG:330900.8:2002JAN18	1527	1654		TM	Cytosolic
61	LG:330900.8:2002JAN18	1655	1672		TM	Transmembrane
61	LG:330900.8:2002JAN18	1673	1677		TM	Extracellular
61	LG:330900.8:2002JAN18	3012	3089	forward 3	SP	
61	LG:330900.8:2002JAN18	3012	3083	forward 3	SP	
61	LG:330900.8:2002JAN18	3012	3089	forward 3	SP	
61	LG:330900.8:2002JAN18	3012	3074	forward 3	SP	
61	LG:330900.8:2002JAN18	3012	3083	forward 3	SP	
61	LG:330900.8:2002JAN18	3012	3077	forward 3	SP	
61	LG:330900.8:2002JAN18	3012	3089	forward 3		
61	LG:330900.8:2002JAN18	3012	3095	forward 3		
61	LG:330900.8:2002JAN18	3012	3080	forward 3	SP	
61	LG:330900.8:2002JAN18	3012	3095	forward 3	SP	
61	LG:330900.8:2002JAN18	3027	3083	forward 3	SP	
62	LG:330931.9:2002JAN18	1	1400		TM	Extracellular
62	LG:330931.9:2002JAN18	1401	1423		TM	Transmembrane
62	LG:330931.9:2002JAN18	1424	1429		TM	Cytosolic
62	LG:330931.9:2002JAN18	1430	1447		TM	Transmembrane
62	LG:330931.9:2002JAN18	1448	1701		TM	Extracellular
62	LG:330931.9:2002JAN18	1702	1724		TM	Transmembrane
62	LG:330931.9:2002JAN18	1725	1816		TM	Cytosolic
62	LG:330931.9:2002JAN18	1817	1839	1	TM	Transmembrane
62	LG:330931.9:2002JAN18	1840	1853	<u> </u>	TM	Extracellular
						Transmembrane
62	LG:330931.9:2002JAN18	1854	1873		TM	Transmembrar

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TABLE 4

		TA	BLE 4						
		Tot	Stop	Fran	ne D	oma	in To	pology	
SEQ ID	Template ID	Start	3100	1.10		уре	-		
NO:			1050			M	(0)	ytosolic	
62	LG:330931.9:2002JAN18	1874	1950			M	Tro	ansmembrane	
62	LG:330931.9:2002JAN18	1951	1973			M	Ex	tracellular	
	LG:330931.9:2002JAN18	1974	1992			ſΜ		ansmembrane	
62	LG:330931.9:2002JAN18	1993	2015			TM		ytosolic	
62	LG:330931.9:2002JAN18	2016	2035			TM	- E	dracellular	
62	LG:330931.9:2002JAN18	1	1109					ansmembrane	
62	LG:330931.9:2002JAN18	1110	1132			TM_		Cytosolic	
62	LG:330931.9:2002JAN18	1133	1144			TM_	- -	ransmembrane	
62	LG:330931.9.200237.116	1145	1167			TM		xtracellular	
62	LG:330931.9:2002JAN18	1168	1701			TM		ransmembrane	
62	LG:330931.9:2002JAN18	1702	1724			TM			\
62	LG:330931.9:2002JAN18	1725	1822			TM		Cytosolic	1
62	LG:330931.9:2002JAN18	1823	1845			TM	7	ransmembrane	1
62	LG:330931.9:2002JAN18	1846	1859			TM		Extracellular	1
62	IG:330931.9:2002JAN18					TM		Transmembrane	-{
62	1G:330931.9:2002JAN18	1860				TM		Cytosolic	4
62	IG:330931.9:2002JAN18	1876				TM		Transmembrane	4
62	IG:330931.9:2002JAN18	1940				TM		Extracellular	4
62	IG:330931.9:2002JAN18	1963				TM		Transmembrane	
62	LG:330931.9:2002JAN18	1997				TM		Cytosolic	
	LG:330931.9:2002JAN18	2020			orward 1				_l
62		266							
62		266			orward 1				7
62		490			orward 1				7
62		490	6 496		orward				_
62		182	6 188	8 f	orward :	2 SP			
62		207	3 218	30 f	forward	3 SP		Cytosolic	一
62			100	5		TN		Transmembran	
63		107	1 129	7		TN		Extracellular	\dashv
6	3 LG:330985.1:2002JAN18			3		TN			<u>, </u>
6	3 LG:330985.1:2002JAN18					TN		Transmembrar	<u>'= </u>
6	3 LG:330985.1:2002JAN18					TN	1	Cytosolic	_
6	3 LG:330985.1:2002JAN18					TI/	Λ	Transmembrar	1e
1	3 I G:330985.1:2002JAN R					TI	V	Extracellular	
	2 IG:330985.1:2002JAN18	3 10				TI	V	Transmembra	ne
	3 I.G:330985.1:2002JAN1	8 19			 	T	VI	Cytosolic	
	53 I G:330985.1:2002JAN1	8 2	'	12			M	Transmembra	ne
	43 I.G:330985.1:2002JAN1	8 22			+		M	Extracellular	
	63 I G:330985.1:2002JAN1	8 24		08	+		M	Transmembro	ne
<u> </u>	63 LG:330985.1:2002JAN1	8 3		31	 		M	Cytosolic	
1	63 LG:330985.1:2002JAN1	8 3		42			M	Transmembro	ne
		8 3		65	+		M	Extracellular	
		18 3	-	69			ſΜ	Transmembro	ane
		18 3		92				Cytosolic	
		18 3	393	98			TM_	Transmembre	ane
	1,0000 IAN			121			TM_	Extracellular	
	- 20000E 1:2002 IAN	10		435			TM_	Transmembr	ane
	63 LG:330985.1:2002JAN	10-		458	1		TM		
	63 LG:330985.1:2002JAN	· <u>· · · ·</u>		470			TM	Cytosolic	
	63 LG:330985.1:2002JAN	•••		490			TM	Transmembr	
	63 LG:330985.1:2002JAN	···	491	1078	1		TM	Extracellular	
<u> </u>	63 LG:330985.1:2002JAN	110	471					- 440	
L									

TABLE 4

			TABLE 4			
SEQID	Template ID	Start	Stop	Frame	15	
NO:		J. G.	Joiop	riame		Topology
63	LG:330985.1:2002JAN18	1798	1857		Туре	
63	LG:330985.1:2002JAN18	1798				
63	LG:330985.1:2002JAN18	1798	- 1.007			
63	LG:330985.1:2002JAN18	1798	1875			
63	LG:330985.1:2002JAN18	550	1872	forward 1	SP	
63	LG:330985.1:2002JAN18		639	forward 1	SP	
63	LG:330985.1:2002JAN18	1798	1863	forward 1	SP	
.63	LG:330985.1:2002JAN18	1798	1842	forward 1	SP	
64	LG:332027.9:2002JAN18	417	476	forward 3	SP	
64	LG:332027.9:2002JAN18	1	570		TM	Extracellular
64	LC:332027.9.2002JAN18	571	593		TM	Transmembrar
	LG:332027.9:2002JAN18	594	613		TM	Cytosolic
64	LG:332027.9:2002JAN18	614	636		TM	Transmembrar
	LG:332027.9:2002JAN18	637	655		TM	Extracellular
	LG:332027.9:2002JAN18	656	678		TM	Transmembrar
	LG:332027.9:2002JAN18	679	865		TM	Cytosolic
	LG:332027.9:2002JAN18	866	888			Transmembran
	LG:332027.9:2002JAN18	889	968		 	Extracellular
64	LG:332027.9:2002JAN18	969	988		 	
64	LG:332027.9:2002JAN18	989	1022		-	Transmembran
64	LG:332027.9:2002JAN18	1	449			Cytosolic
64	LG:332027.9:2002JAN18	450	472			Extracellular
_64	LG:332027.9:2002JAN18	473	553			Transmembran
64	LG:332027.9:2002JAN18	554	576			Cytosolic
64	G:332027.9:2002JAN18	577	624	- -		Transmembrane
	G:332027.9:2002JAN18	625	644			Extracellular
64 L	G:332027.9:2002JAN18	645	859			Transmembrane
64 L	.G:332027.9:2002JAN18	860	882			Cytosolic
64 L	.G:332027.9:2002JAN18	883	918		TM 1	ransmembrane
64 L	G:332027.9:2002JAN18	919				xtracellular
64 L	G:332027.9:2002JAN18	942	941		TM T	ransmembrane
64 L	G:332027.9:2002JAN18	962	961		TM (Cytosolic
'64 L	G:332027.9:2002JAN18		984		T M	ransmembrane
64 L	G:332027.9:2002JAN18	985	1021			xtracellular
'64 L	G:332027.9:2002JAN18	757	750		M E	xtracellular
	G:332027.9:2002JAN18	751	773		M T	ransmembrane
64 LC	G:332027.9:2002JAN18	774	800			ytosolic
	G:332027.9:2002JAN18	801	823		M Tr	ansmembrane
	G:332027.9:2002JAN18	824	872	T	M E	xtracellular
	9:332027.9:2002JAN18	873	895	T		ansmembrane
	9:332027.9:2002JAN18 9:332027.9:2002JAN18	896	948	T	м с	ytosolic
	2:332027 0:2000 (ANTO	949	971			ansmembrane
	9:332027.9:2002JAN18	972	1021			dracellular
	9:335377.8:2002JAN18	3010	3108	forward 1 S		
	9:335377.8:2002JAN18	3010	3084	forward 1 S		
	9:335377.8:2002JAN18	2090	2143	forward 2 SI		
	9:335377.8:2002JAN18	2883	2945	forward 3 SI		
65 LG	9:335377.8:2002JAN18	2883	2957	forward 3 SI		
65 LG	5:335377.8:2002JAN18	2883	2951	forward 3 SF		
65 LG	335377.8:2002JAN18 335377.8:2002JAN18	3030	3110	forward 3 SF		
65 LG						

TABLE 4

SEQ ID	Template ID	Start	Stop	Frame	Domain	Topology
NO:		Joidin	CiOp	Traine	Туре	lopolog,
66	LG:337452.25:2002JAN18	1156	1233	forward 1	SP	
66	LG:337452.25:2002JAN18	2237	2314	forward 2	SP	
66	LG:337452.25:2002JAN18	1514	1600	forward 2	SP	
67	LG:340580.16:2002JAN18	1	393	101Wara 2	TM	Extracellular
67	LG:340580.16:2002JAN18	394	416		TM	Transmembrane
67	LG:340580.16:2002JAN18	417	422		TM	Cytosolic
67	LG:340580.16:2002JAN18	423	440		TM	Transmembrane
67	LG:340580.16:2002JAN18	441	1131		TM	Extracellular
67	LG:340580.16:2002JAN18	1132	1151		TM	Transmembrane
67	LG:340580.16:2002JAN18	1152	1685		TM	Cytosolic
67	LG:340580.16:2002JAN18	1686	1708		TM	Transmembrane
67	LG:340580.16:2002JAN18	1709	1722		TM	Extracellular
67	LG:340580.16:2002JAN18	1723	1745		TM	Transmembrane
67	LG:340580.16:2002JAN18	1746	1796	-	TM	Cytosolic
67	LG:340580.16:2002JAN18	1797	1819		TM	Transmembrane
67	LG:340580.16:2002JAN18	1820	1881		TM	Extracellular
67	LG:340580.16:2002JAN18	1882	1901		TM	Transmembrane
67	LG:340580.16:2002JAN18	1902	1975		TM	Cytosolic
67	LG:340580.16:2002JAN18	1976	1998		TM	Transmembrane
67	LG:340580.16:2002JAN18	1999	2288		TM	Extracellular
67	LG:340580.16:2002JAN18	2289	2311		TM	Transmembrane
67	LG:340580.16:2002JAN18	2312	2317	-	TM	Cytosolic
67	LG:340580.16:2002JAN18	2318	2335		TM	Transmembrane
67	LG:340580.16:2002JAN18	2336	2446	 	TM	Extracellular
67	LG:340580.16:2002JAN18	1	. 42		TM	Cytosolic
67	LG:340580.16:2002JAN18	43	61	-	TM	Transmembrane
67	LG:340580.16:2002JAN18	62	80		TM	Extracellular
67	LG:340580.16:2002JAN18	81	98		TM	Transmembrane
67	LG:340580.16:2002JAN18	99	197		TM	Cytosolic
67	LG:340580.16:2002JAN18	198	220		TM	Transmembrane
67	LG:340580.16:2002JAN18	221	393		TM	Extracellular
67	LG:340580.16:2002JAN18	394	416		TM	Transmembrane
67	LG:340580.16:2002JAN18	417	428		TM	Cytosolic
67	LG:340580.16:2002JAN18	429	451		TM	Transmembrane
67	LG:340580.16:2002JAN18	452	2288		TM	Extracellular
67	LG:340580.16:2002JAN18	2289	2311		TM	Transmembrane
67	LG:340580.16:2002JAN18	2312	2323		TM	Cytosolic
67	LG:340580.16:2002JAN18	2324	2346		TM	Transmembrane
67	LG:340580.16:2002JAN18	2347	2400		TM	Extracellular
67	LG:340580.16:2002JAN18	2401	2423		TM	Transmembrane
67	LG:340580.16:2002JAN18	2424	2446		TM	Cytosolic
67	LG:340580.16:2002JAN18	1	80		TM	Cytosolic
67	LG:340580.16:2002JAN18	81	103		TM	Transmembrane
67	LG:340580.16:2002JAN18	104	391		TM	Extracellular
67	LG:340580.16:2002JAN18	392	414		TM	Transmembrane
67	LG:340580.16:2002JAN18	415	426		TM	Cytosolic
67	LG:340580.16:2002JAN18	427	449		TM	Transmembrane
67	LG:340580.16:2002JAN18	450	1729		TM	Extracellular
67	LG:340580.16:2002JAN18	1730	1761		TM	Transmembrane

TABLE 4

SEQ ID No:	
67 LG:340580.16:2002JAN18 1762 1791 TM Cytosolic 67 LG:340580.16:2002JAN18 1792 1814 TM Transmer 67 LG:340580.16:2002JAN18 1815 1835 TM Transmer 67 LG:340580.16:2002JAN18 1836 1853 TM Transmer 67 LG:340580.16:2002JAN18 1854 1873 TM Cytosolic 67 LG:340580.16:2002JAN18 1897 1979 TM Transmer 67 LG:340580.16:2002JAN18 1980 2002 TM Transmer 67 LG:340580.16:2002JAN18 2003 2092 TM Cytosolic 67 LG:340580.16:2002JAN18 2003 2015 TM Transmer 67 LG:340580.16:2002JAN18 2003 2115 TM Transmer 67 LG:340580.16:2002JAN18 2310 2321 TM Cytosolic 67 LG:340580.16:2002JAN18 2310 2322 344 TM Transmer	
67 LG:340580.16:2002JAN18 1815 1835 TM Extracello Cytosolic Cig. 340580.16:2002JAN18 1815 1835 TM Extracello Cytosolic Cig. 340580.16:2002JAN18 1854 1873 TM Cytosolic Cytosolic Cig. 340580.16:2002JAN18 1854 1873 TM Cytosolic Cig. 340580.16:2002JAN18 1854 1873 TM Cytosolic Cig. 340580.16:2002JAN18 1854 1873 TM Cytosolic Cig. 340580.16:2002JAN18 1897 1979 TM Extracello Cig. 340580.16:2002JAN18 1897 1979 TM Extracello Cig. 340580.16:2002JAN18 1898 2002 TM Transmer Cig. 340580.16:2002JAN18 1898 2002 TM Cytosolic Cig. 340580.16:2002JAN18 2003 2092 TM Cytosolic Cig. 340580.16:2002JAN18 2003 2092 TM Cytosolic Cig. 340580.16:2002JAN18 2003 2092 TM Extracello Cig. 340580.16:2002JAN18 2003 2092 TM Extracello Cig. 340580.16:2002JAN18 2003 2092 TM Extracello Cig. 340580.16:2002JAN18 2116 2286 TM Extracello Cig. 340580.16:2002JAN18 2116 2286 TM Extracello Cig. 340580.16:2002JAN18 2310 2321 TM Cytosolic Cig. 340580.16:2002JAN18 2310 2321 TM Cytosolic Cig. 340580.16:2002JAN18 2310 2321 TM Cytosolic Cig. 340580.16:2002JAN18 2345 2363 TM Extracello Cig. 340580.16:2002JAN18 2345 2363 TM Cig. 340580.16:2002JAN18 2345 2363 TM Cig. 340580.16:2002JAN18 2345 2363 Tig. 2415 TM Cig. 340580.16:2002JAN18 2345 2345 Tig. 2446 TM Transmer Cig. 340580.16:2002JAN18 2345 2345 Torward 1 SP Cig. 340580.16:2002JAN18 2345 2345 Torward 2 SP Cig. 340580.16:2002JAN18 2345 2345 Torward 2 SP Cig. 340580.16:2002JAN18 2573 2656 Torward 2 SP Cig. 340580.16:2002JAN18 2573 2656 Torward 2 SP Cig. 340580.16:2002JAN18 2573 2656 Torward 3 SP Cig. 340580.16:2002JAN18 2573 2656 Torward 3 SP Cig. 340580.16:2002JAN18 2573 2656 Torward 3 SP Cig. 340580.16:2002JAN18 2573 2656 Torwar	
67 LG:340580.16:2002JAN18 1836 1853 TM Extracellic G7 LG:340580.16:2002JAN18 1836 1853 TM Transmer G7 LG:340580.16:2002JAN18 1854 1873 TM Cytosolic G7 LG:340580.16:2002JAN18 1874 1896 TM Transmer G7 LG:340580.16:2002JAN18 1897 1979 TM Extracellic G7 LG:340580.16:2002JAN18 1897 1979 TM Extracellic G7 LG:340580.16:2002JAN18 1980 2002 TM Transmer G7 LG:340580.16:2002JAN18 1980 2002 TM Cytosolic G7 LG:340580.16:2002JAN18 2003 2092 TM Cytosolic G7 LG:340580.16:2002JAN18 2003 2092 TM Cytosolic G7 LG:340580.16:2002JAN18 2003 2092 TM Transmer G7 LG:340580.16:2002JAN18 2093 2115 TM Transmer G7 LG:340580.16:2002JAN18 2116 2286 TM Extracellic G7 LG:340580.16:2002JAN18 2116 2286 TM Extracellic G7 LG:340580.16:2002JAN18 2310 2321 TM Cytosolic G7 LG:340580.16:2002JAN18 2310 2321 TM Cytosolic G7 LG:340580.16:2002JAN18 2332 2344 TM Transmer G7 LG:340580.16:2002JAN18 2345 2363 TM Extracellic G7 LG:340580.16:2002JAN18 2345 2363 TM Extracellic G7 LG:340580.16:2002JAN18 2345 2363 TM Extracellic G7 LG:340580.16:2002JAN18 2364 2386 TM Extracellic G7 LG:340580.16:2002JAN18 2365 2565 5973 TM Cytosolic G7 LG:340580.16:2002JAN18 5926 5985 Torward 1 SP C7 LG:340580.16:2002JAN18 5926 5985 Torward 2 SP C7 LG:340580.16:2002JAN18 5926 5985 Torward 3 SP C7 LG:340580.16:2002JAN18 5926 5985 Torward 3 SP C7 LG:340580.16:2002JAN18 5926 5985 Torward 3 SP C7 LG:340580.16:2002JAN18 5987 5258 Torward 3 SP C7 LG:34	
67 LG:340580.16:2002JAN18 1836 1853 TM Transmer 67 LG:340580.16:2002JAN18 1854 1873 TM Cytosolic 7 LG:340580.16:2002JAN18 1874 1896 TM Transmer 67 LG:340580.16:2002JAN18 1897 1979 TM Extracellt 67 LG:340580.16:2002JAN18 1980 2002 TM Transmer 67 LG:340580.16:2002JAN18 1980 2002 TM Transmer 67 LG:340580.16:2002JAN18 1980 2002 TM Cytosolic 7 LG:340580.16:2002JAN18 1980 2002 TM Transmer 67 LG:340580.16:2002JAN18 2003 2092 TM Cytosolic 7 LG:340580.16:2002JAN18 2003 2092 TM Cytosolic 7 LG:340580.16:2002JAN18 2093 2115 TM Transmer 67 LG:340580.16:2002JAN18 2116 2286 TM Extracellt 67 LG:340580.16:2002JAN18 2116 2286 TM Cytosolic 7 LG:340580.16:2002JAN18 2310 2321 TM Cytosolic 7 LG:340580.16:2002JAN18 2310 2321 TM Cytosolic 67 LG:340580.16:2002JAN18 2310 2321 TM Cytosolic 7 LG:340580.16:2002JAN18 2345 2363 TM Extracellt 67 LG:340580.16:2002JAN18 2345 2363 TM Extracellt 67 LG:340580.16:2002JAN18 2345 2363 TM Extracellt 67 LG:340580.16:2002JAN18 2387 2392 TM Cytosolic 7 LG:340580.16:2002JAN18 2387 2392 TM Cytosolic 67 LG:340580.16:2002JAN18 2387 2392 TM Cytosolic 67 LG:340580.16:2002JAN18 2393 2415 TM Transmer 67 LG:340580.16:2002JAN18 5926 5973 forward 1 SP 67 LG:340580.16:2002JAN18 5926 5985 forward 1 SP 67 LG:340580.16:2002JAN18 5926 5985 forward 2 SP 67 LG:340580.16:2002JAN18 5926 6000 forward 1 SP 67 LG:340580.16:2002JAN18 5926 6000 forward 1 SP 67 LG:340580.16:2002JAN18 5926 6000 forward 2 SP 67 LG:340580.16:2002JAN18 5926 forward 2 SP 67 LG:340580.16:2002JAN18 5926 forward 2 SP 67 LG:340580.16:2002JAN18 5926 forward 2 SP 67 LG:340580.16:2002JAN18 5187 5264 forward 3 SP 67 LG:340580.16:2002JAN18 5187 5264 forward 3 SP 67 LG:340580.16:2002JAN18 5187 5264 forward 3 SP 67 LG:340580.16:2002JAN18 5187 5265 forward 3 SP 67 LG:340580.16:2002JAN18 5187 5265 forward 3 SP 67 LG:340580.16:2002JAN18 5187 5265 forward 3 SP 67 LG:340580.16:2	
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68 LG:350272.6:2002JAN18 673 679 TM Cytosoli 68 LG:350272.6:2002JAN18 1967 2038 forward 2 SP	mbrane
68 LG:350272.6:2002JAN18 1967 2038 forward 2 SP	<u> </u>
00 100.000272:0:20020711110	
68 LG:350272.6:2002JAN18 1967 2032 forward 2 SP	
68 LG:350272.6:2002JAN18 1967 2038 forward 2 SP	
68 LG:350272.6:2002JAN18 1967 2035 forward 2 SP	
68 LG:350272.6:2002JAN18 1793 1891 forward 2 SP	
68 LG:350272.6:2002JAN18 1967 2026 forward 2 SP	
68 LG:350272.6:2002JAN18 1787 1885 forward 2 SP	
68 LG:350272.6:2002JAN18 1218 1277 forward 3 SP	
68 LG:350272.6:2002JAN18 1218 1274 forward 3 SP	
68 LG:350272.6:2002JAN18 1218 1280 forward 3 SP	
69 LG:397228.1:2002JAN18 1 106 TM Cytosol	C
70 LG:401325.41:2002JAN18 1867 1959 forward 1 SP	

TABLE 4

	<u></u>	17 (DLC 4			
SEQ ID	Template ID	Start	Stop	Frame	i	Topology
NO:			ļ <u>.</u>	ļ	Туре	
70	LG:401325.41:2002JAN18	2546	2617	forward 2	SP	
70	LG:401325.41:2002JAN18	2546	2611	forward 2	SP	
70	LG:401325.41:2002JAN18	2546	2617	forward 2	SP	
70	LG:401325.41:2002JAN18	2546	2596	forward 2	SP	
70	LG:401325.41:2002JAN18	2666	2722	forward 2	SP	
71	LG:402029.14:2002JAN18	1	207		TM	Extracellular
71	LG:402029.14:2002JAN18	208	230		TM	Transmembrane
71	LG:402029.14:2002JAN18	231	305		TM	Cytosolic
71	LG:402029.14:2002JAN18	306	325		TM	Transmembrane
71	LG:402029.14:2002JAN18	326	339		TM	Extracellular
71	LG:402029.14:2002JAN18	340	357		TM	Transmembrane
71	LG:402029.14:2002JAN18	358	369		TM	Cytosolic
71	LG:402029.14:2002JAN18	370	392		TM	Transmembrane
71	LG:402029.14:2002JAN18	393	1147		TM	Extracellular
71	LG:402029.14:2002JAN18	2527	2607	forward 1	SP	
71	LG:402029.14:2002JAN18	1342	1431	forward 1	SP	
72	LG:407233.2:2002JAN18	1	773		TM	Extracellular
72	LG:407233.2:2002JAN18	774	796		TM	Transmembrane
72	LG:407233.2:2002JAN18	797	798		TM	Cytosolic
72	LG:407233.2:2002JAN18	1	57		TM	Cytosolic
72	LG:407233.2:2002JAN18	58	80		TM	Transmembrane
72	LG:407233.2:2002JAN18	81	798		TM	Extracellular
72	LG:407233.2:2002JAN18	1	50		TM	Extracellular
72	LG:407233.2:2002JAN18	51	73		TM	Transmembrane
72	LG:407233.2:2002JAN18	74	390		TM	Cytosolic
72	LG:407233.2:2002JAN18	391	413		TM	Transmembrane
72	LG:407233.2:2002JAN18	414	689		TM	Extracellular
72	LG:407233.2:2002JAN18	690	712		TM	Transmembrane
1	LG:407233.2:2002JAN18	713	798		TM	Cytosolic
72	LG:407233.2:2002JAN18	1180	1266	forward 1		
72	LG:407233.2:2002JAN18	139	213	forward 1	SP	
72	LG:407233.2:2002JAN18	1192	1257	forward 1		
72		1204	1266	forward 1		
72	LG:407233.2:2002JAN18	1182	1265	forward 3		
72	LG:407233.2:2002JAN18	1 102	519	lorwara	TM	Extracellular
73	LG:407346.1:2002JAN18	520	542		TM	Transmembrane
73	LG:407346.1:2002JAN18		562		TM	Cytosolic
73	LG:407346.1:2002JAN18	543			TM	Transmembrane
73	LG:407346.1:2002JAN18	563	585 1725		TM	Extracellular
73	LG:407346.1:2002JAN18	586	1748		TM	Transmembrane
73	LG:407346.1:2002JAN18	1726	2071		TM	Cytosolic
73	LG:407346.1:2002JAN18	1749			TM	Transmembrane
73	LG:407346.1:2002JAN18	2072	2094		TM	Extracellular
73	LG:407346.1:2002JAN18	2095	2126		TM	Transmembrane
73	LG:407346.1:2002JAN18	2127	2149		TM	Cytosolic
73	LG:407346.1:2002JAN18	2150	2161			Extracellular
73	LG:407346.1:2002JAN18	1	2001		TM	Transmembrane
73	LG:407346.1:2002JAN18	2002	2021		TM	Cytosolic
73	LG:407346.1:2002JAN18	2022	2161		TM	Extracellular
73	LG:407346.1:2002JAN18]]	1616		TM	EXITACEIMICI

TABLE 4

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SEQ ID	Template ID	Start	Stop	Frame	Domain	Topology
NO:	1				Туре	, ,
73	LG:407346.1:2002JAN18	1617	1639		TM	Transmembrane
73	LG:407346.1:2002JAN18	1640	1741		TM	Cytosolic
73	LG:407346.1:2002JAN18	1742	1764		TM	Transmembrane
73	LG:407346.1:2002JAN18	1765	2161		TM	Extracellular
74	LG:407689.7:2002JAN18	1	2173		TM	Extracellular
74	LG:407689.7:2002JAN18	2174	2196		TM	Transmembrane
74	LG:407689.7:2002JAN18	2197	2202		TM	Cytosolic
74	LG:407689.7:2002JAN18	2203	2225		TM	Transmembrane
74	LG:407689.7:2002JAN18	2226	2259		TM	Extracellular
74	LG:407689.7:2002JAN18	2188	2262	forward 1	SP	
74	LG:407689.7:2002JAN18	4693	4752	forward 1	SP	
74	LG:407689.7:2002JAN18	2182	2262	forward 1	SP	
74	LG:407689.7:2002JAN18	2104	2181	forward 1	SP	
74	LG:407689.7:2002JAN18	1378	1458	forward 1	SP	
74	LG:407689.7:2002JAN18	2023	2094	forward 1	SP	
74	LG:407689.7:2002JAN18	4693	4755	forward 1	SP	
74	LG:407689.7:2002JAN18	6269	6331	forward 2	SP	
74	LG:407689.7:2002JAN18	1206	1274	forward 3	SP	
75	LG:407700.1:2002JAN18	1	20		TM	Cytosolic
75	LG:407700.1:2002JAN18	21	43		TM	Transmembrane
75	LG:407700.1:2002JAN18	44	750		TM	Extracellular
75	LG:407700.1:2002JAN18	1	571		TM	Extracellular
75	LG:407700.1:2002JAN18	572	594		TM	Transmembrane
75	LG:407700.1:2002JAN18	595	714		TM	Cytosolic
75	LG:407700.1:2002JAN18	715	737		TM	Transmembrane
75	LG:407700.1:2002JAN18	738	750		TM	Extracellular
.75	LG:407700.1:2002JAN18	47	106	forward 2	SP	
75	LG:407700.1:2002JAN18	47	106	forward 2	SP	
75	LG:407700.1:2002JAN18	47	139	forward 2	SP	
75	LG:407700.1:2002JAN18	47	103	forward 2	SP	
75	LG:407700.1:2002JAN18	47	109	forward 2	SP	
75	LG:407700.1:2002JAN18	378	464	forward 3	SP	
76	LG:410461.92:2002JAN18	1	262		TM	Extracellular
76	LG:410461.92:2002JAN18	263	285		TM	Transmembrane
76	LG:410461.92:2002JAN18	286	492		TM	Cytosolic
76	LG:410461.92:2002JAN18	493	515		TM	Transmembrane
76	LG:410461.92:2002JAN18	516	950		TM	Extracellular
76	LG:410461.92:2002JAN18	258	344	forward 3	SP	
76	LG:410461.92:2002JAN18	258	344	forward 3	SP	
76	LG:410461.92:2002JAN18	258	338	forward 3	SP	
76	LG:410461.92:2002JAN18	273	344	forward 3	SP	
76	LG:410461.92:2002JAN18	273	338	forward 3	SP	
76	LG:410461.92:2002JAN18	258	338	forward 3	SP	
76	LG:410461.92:2002JAN18	276	338	forward 3	SP	
77	LG:411043.3:2002JAN18	1	121		TM	Cytosolic
77	LG:411043.3:2002JAN18	122	144		TM	Transmembrane
77	LG:411043.3:2002JAN18	145	203	_	TM	Extracellular
77	LG:411043.3:2002JAN18	204	223		TM	Transmembrane
77	LG:411043.3:2002JAN18	224	458		TM	Cytosolic

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SEQ ID	Template ID	Start	Stop	Frame	:	Topology
NO:			100		Туре	T
77	LG:411043.3:2002JAN18	459	481		TM	Transmembrane
77	LG:411043.3:2002JAN18	482	707	 	TM	Extracellular
77	LG:411043.3:2002JAN18	606	668	forward 3	SP	
78	LG:438690.47:2002JAN18	1	1512	_	TM	Extracellular
78	LG:438690.47:2002JAN18	1513	1535		TM	Transmembrane
78	LG:438690.47:2002JAN18	1536	1767		TM	Cytosolic
78	LG:438690.47:2002JAN18	1768	1787		TM	Transmembrane
78	LG:438690.47:2002JAN18	1788	1788		TM	Extracellular
78	LG:438690.47:2002JAN18	1	1506		TM	Extracellular
78	LG:438690.47:2002JAN18	1507	1529		TM	Transmembrane
78	LG:438690.47:2002JAN18	1530	1662		TM	Cytosolic
78	LG:438690.47:2002JAN18	1663	1685		TM	Transmembrane
78	LG:438690.47:2002JAN18	1686	1788		TM	Extracellular
78	LG:438690.47:2002JAN18	3451	3522	forward 1	SP	
78	LG:438690.47:2002JAN18	1777	1866	forward 1	SP	
78	LG:438690.47:2002JAN18	3451	3522	forward 1	SP	
78	LG:438690.47:2002JAN18	3451	3525	forward 1	SP	
78	LG:438690.47:2002JAN18	3032	3085	forward 2	SP	
78	LG:438690.47:2002JAN18	3032	3091	forward 2	SP	
78	LG:438690.47:2002JAN18	1043	1120	forward 2	SP	
78	LG:438690.47:2002JAN18	3032	3100	forward 2	SP	
78	LG:438690.47:2002JAN18	582	650	forward 3	SP	
79	LG:444677.81:2002JAN18	1055	1111	forward 2	SP	
79	LG:444677.81:2002JAN18	818	880	forward 2	SP	
79	LG:444677.81:2002JAN18	234	· 293	forward 3	SP	
80	LG:457464.24:2002JAN18	1	438		TM	Extracellular
80	LG:457464.24:2002JAN18	439	461		TM	Transmembrane
80	LG:457464.24:2002JAN18	462	536		TM	Cytosolic
80	LG:457464.24:2002JAN18	1000	1092	forward 1	SP	
80	LG:457464.24:2002JAN18	1347	1400	forward 3	SP	
80	LG:457464.24:2002JAN18	1347	1403	forward 3	SP	
81	LG:7684793.15:2002JAN18	940	1002	forward 1	SP	
81	LG:7684793.15:2002JAN18	940	1008	forward 1	SP	
81	LG:7684793.15:2002JAN18	3253	3345	forward 1	SP	
81	LG:7684793.15:2002JAN18	3155	3244	forward 2	SP	
81	LG:7684793.15:2002JAN18	2117	2194	forward 2	SP	
81	LG:7684793.15:2002JAN18	2114	2188	forward 2	SP	
81	LG:7684793.15:2002JAN18	3984	4058	forward 3		
82	LG:7687485.1:2002JAN18	1	252		TM	Cytosolic
82	LG:7687485.1:2002JAN18	253	275		TM	Transmembrane
82	LG:7687485.1:2002JAN18	276	380		TM	Extracellular
82	LG:7687485.1:2002JAN18	781	828	forward 1	SP	
82	LG:7687485.1:2002JAN18	781	834	forward 1		
82	LG:7687485.1:2002JAN18	299	367	forward 2		
82	LG:7687485.1:2002JAN18	299	361	forward 2		
83	LG:7689661.4:2002JAN18	1	345		TM	Extracellular
83	LG:7689661.4:2002JAN18	346	365		TM	Transmembrane
83	LG:7689661.4:2002JAN18	366	509		TM	Cytosolic
83	LG:7689661.4:2002JAN18	510	532		TM	Transmembrane

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		TAB	_E 4			
	TInto ID	Start	Stop	Frame	Domain	Topology
	Template ID	Joidin	O.O.D		Туре	
NO:	1. 7. 7.00((1. 4.0000 IANI)	533	757		TM	Extracellular
83	LG:7689661.4:2002JAN18	1562	1618	forward 2	SP	
83	LG:7689661.4:2002JAN18	1	95	101111111	TM	Cytosolic
84	LG:7690373.1:2002JAN18	96	118		TM	Transmembrane
84	LG:7690373.1:2002JAN18	119	219		TM	Extracellular
84	LG:7690373.1:2002JAN18	119	556	 	TM	Extracellular
85	LG:7696560.1:2002JAN18	557	576		TM	Transmembrane
85	LG:7696560.1:2002JAN18	577	602	1	TM	Cytosolic
85	LG:7696560.1:2002JAN18	1135	1194	forward 1	SP	
85	LG:7696560.1:2002JAN18	83	166	forward 2	SP	
85	LG:7696560.1:2002JAN18		1466	101Waru 2	TM	Extracellular
86	LG:7698190.26:2002JAN18	1 147	1489		TM	Transmembrane
86	LG:7698190.26:2002JAN18	1467	1521	 	TM	Cytosolic
86	LG:7698190.26:2002JAN18	1490		 	TM	Extracellular
86	LG:7698190.26:2002JAN18	1	1460	 	TM	Transmembrane
86	LG:7698190.26:2002JAN18	1461	1483		TM	Cytosolic
86	LG:7698190.26:2002JAN18	1484	1521		TM	Extracellular
86	LG:7698190.26:2002JAN18	11	1465		TM	Transmembrane
86	LG:7698190.26:2002JAN18	1466	1488		TM	Cytosolic
86	LG:7698190.26:2002JAN18	1489	1520	forward 1	SP	0710000
86	LG:7698190.26:2002JAN18	2872	2976	forward 1		· · · · · · · · · · · · · · · · · · ·
86	LG:7698190.26:2002JAN18	4408	4470	forward 1		
87	LG:7763560.12:2002JAN18	1162	1260	forward 2		
87	LG:7763560.12:2002JAN18	2225	2308	forward 2		
87	LG:7763560.12:2002JAN18	2225	2308	forward 2		
88	LG:7763587.20:2002JAN18	431	523	forward 2		
88	LG:7763587.20:2002JAN18	476	520	forward 2		
88	LG:7763587.20:2002JAN18	476	550			
88	LG:7763587.20:2002JAN18	1692	1787	forward :	TM	Extracellular
89	LG:899263.10:2002JAN18	1	784		TM	Transmembrane
89	LG:899263.10:2002JAN18	785	807			Cytosolic
89	LG:899263.10:2002JAN18	808	911	6	TM 3 SP	Cytosolic
89	LG:899263.10:2002JAN18	93	167	forward		Extracellular
90	LG:977837.31:2002JAN18	1	55		TM	Transmembrane
90	LG:977837.31:2002JAN18	56	78			Cytosolic
90	LG:977837.31:2002JAN18	79	84		TM	Transmembrane
90	LG:977837.31:2002JAN18	85	107		TM	Extracellular
90	LG:977837.31:2002JAN18	108	320		TM	Extracellular
91	LG:978560.13:2002JAN18		1399		TM	Transmembrane
91	LG:978560.13:2002JAN18	1400	1419		TM	Cytosolic
91	LG:978560.13:2002JAN18	1420	1426		TM	Extracellular
91	LG:978560.13:2002JAN18	1	1396		TM	Transmembrane
91	LG:978560.13:2002JAN18	1397	1419		TM	Cytosolic
91	LG:978560.13:2002JAN18	1420	1425		TM	Cytosolic
91	LG:978560.13:2002JAN18	2735	2806	forward		
91	LG:978560.13:2002JAN18	3971	4039	forward		
91		3971	4033	forward		
91		2735				
91	LG:978560.13:2002JAN18	2735				
, , ,	LG:978560.13:2002JAN18	3971	4030	forward	12 ISP	1

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TABLE 4

		TABL	.E 4			
		Start	Stop	Frame	Domain	Topology
SEQ ID	Template ID	Sidii	Siop	Truine	Туре	
NO:		2073	4033	forward 2	SP	
91	LG:978560.13:2002JAN18	3971 3621	3677	forward 3	SP	
91	LG:978560.13:2002JAN18	3621	3665	forward 3	SP	
91	LG:978560.13:2002JAN18		650	forward 3	SP	
91	LG:978560.13:2002JAN18	579	650	forward 3	SP	
91	LG:978560.13:2002JAN18	558	656	forward 3	SP	
91	LG:978560.13:2002JAN18	579	656	forward 3	SP	
91	LG:978560.13:2002JAN18	558	656	forward 3		
91	LG:978560.13:2002JAN18	558	948	forward 1	SP	
92	LG:979390.2:2002JAN18	886	938	forward 3		
92	LG:979390.2:2002JAN18	861	605	forward 3		
92	LG:979390.2:2002JAN18	543	953	forward 3		
92	LG:979390.2:2002JAN18	861	614	forward 3		
92	LG:979390.2:2002JAN18	543	1397	forward 3		
93	LG:983019.1:2002JAN18	1338	1795	10. Wala	TM	Extracellular
94	LG:997202.7:2002JAN18	1	1818		TM	Transmembrane
94	LG:997202.7:2002JAN18	1796	1837		TM	Cytosolic
94	LG:997202.7:2002JAN18	1819	1860		TM	Transmembrane
94	LG:997202.7:2002JAN18	1838	1869		TM	Extracellular
94	LG:997202.7:2002JAN18	1861			TM	Transmembrane
94	LG:997202.7:2002JAN18	1870	1892		TM	Cytosolic
94	LG:997202.7:2002JAN18	1893	2198	forward		
94	LG:997202.7:2002JAN18	3064	3162	forward		
94	LG:997202.7:2002JAN18	3076	3153	forward		
94	LG:997202.7:2002JAN18	4567	4626	forward		
94	LG:997202.7:2002JAN18	4024	4095	forward		
94	LG:997202.7:2002JAN18	3086	3148	forward		
94	LG:997202.7:2002JAN18	3086	3157	forward		
94	LG:997202.7:2002JAN18	3086	3154	forward		
94	LG:997202.7:2002JAN18	3086	3163	forward		
94	LG:997202.7:2002JAN18	3086	3163	forward		
94	LG:997202.7:2002JAN18	6288	6332	forward		
94	LG:997202.7:2002JAN18	738	794	forward		
94	LG:997202.7:2002JAN18	6288	6347	forward		
94	LG:997202.7:2002JAN18	6288	6347	forward		
94	LG:997202.7:2002JAN18	1140	1214	forward		
94	LG:997202.7:2002JAN18	1467				
94	LG:997202.7:2002JAN18	1134				
9/	LG:997202.7:2002JAN18	5583		lorwan	TM	Cytosolic
95	LG:998756.3:2002JAN18		57		TM	Transmembrane
98	LG:998756.3:2002JAN18	58	77		TM	Extracellular
9:	5 LG:998756.3:2002JAN18	78	80		TM	Transmembrane
9	5 LG:998756.3:2002JAN18	81	103		TM	Cytosolic
9	5 LG:998756.3:2002JAN18	104	208		TM	Transmembrane
9	5 LG:998756,3:2002JAN18	209	231		TM	Extracellular
	5 LG:998756.3:2002JAN18	232	245		TM	Transmembrane
	5 LG:998756.3:2002JAN18	240	268		TM	
	5 LG:998756.3:2002JAN18	269			TM	
	5 LG:998756.3:2002JAN18	298			TM	
I	D5 LG:998756.3:2002JAN18	321	145	<u> </u>	11.01	

TABLE 4

ID Template ID			ABLE 4			
ID Template ID		Start	Stop	Frame	Doma	in Topology
			_		Туре	"I Topology
LG:998756.3:	2002JAN18	1	175		TM	Extracellular
LG:998756.3:	2002JAN18	176	198		TM	
LG:998756.3:	2002JAN18	199	204		TM	Transmembrar
LG:998756.3:	2002JAN18	205	224		TM	Cytosolic
LG:998756.3:	81/AL2002	225	1455			Transmembrar
LG:998756.3:	2002JAN18	157	234	forward	TM	Extracellular
LG:998756.3:	2002JAN18	157	237			
LG:998756.3:	2002JAN18	157	216	forward		
LG:998756.3:	2002JAN18	157	234	forward		
LG:998756.3:2	2002.JAN18	716	790	forward		
LG:998756.3:2		716		forward 2		
LG:998756.3:2	002371118		796	forward 2		
LG:998756.3:2	002071110	716	796	forward 2		
LG:998756.3:2	0023/41116	716	799	forward 2		
LG:998756.3:2		1461	1553	forward 3	SP SP	
LG:103460.00	002JAN18	1461	1559	forward 3	SP	
LG:103460.28	2002JAN18	1	88		TM	Cytosolic
LG:103460.28	2002JAN18	89	111		TM	Transmembrane
LG:103460.28:	2002JAN18	112	500		TM	Extracellular
LG:103460.28:	2002JAN18	501	523		TM	Transmembrane
LG:103460.28:	2002JAN18	524	663		TM.	
LG:103460.28:	2002JAN18	1	87		TM	Cytosolic
LG:103460.28:	2002JAN18	88	105		TM	Cytosolic
LG:103460.28:	2002JAN18	106	662			Transmembrane
LG:103460.28:	2002JAN18	256	324	forward 1	TM	Extracellular
LG:103460.28:	2002JAN18	256	339		SP	
LG:103460.28:	2002JAN18	256	330	forward 1	SP	
LG:103460.28:2	2002JAN18	256	330	forward 1	SP	
LG:103460.28:2	002JAN18	256	324	forward 1	SP	
LG:103460.28:2	002 JAN18	1029		forward 1	SP	
LG:1501505.19		1029	1100	forward 3	SP	
LG:1501505.19	20023/1110	1	48		TM	Cytosolic
LG:1501505.19	2002JAN10	49	71			Transmembrane
LG:1501505.19:	2002JAN18	72	80		TM	Extracellular
LC:1501505.19	2002JAN18	81	103			Transmembrane
LG:1501505.19:	2002JAN 18	104	109			Cytosolic
LG:1501505.19:	2002JAN18	110	132			Transmembrane
LG:1501505.19:	2002JAN18	133	368			Extracellular
LG:233444.9:20	J2JAN18	1	33			Cytosolic
LG:233444.9:20	JZJAN18	34	56			Transmembrane
LG:233444.9:20	DZJAN18	57	681			xtracellular
LG:233444.9:20)2JAN18	682	704			
LG:233444.9:200)2JAN18	705	723			ransmembrane
LG:233444.9:200)2JAN18	724	746			Cytosolic
LG:233444.9:200	2JAN18	747	911			ransmembrane
LG:233444.9:200	2JAN18	1	69			xtracellular
LG:233444.9:200	2JAN18	70	89			Cytosolic
LG:233444.9:200	2JAN18	90	103			ransmembrane
LG:233444.9:200		104				xtracellular
LG:233444.9:200	2.IAN18		126			ransmembrane
LG:233444 0:200	2 14 110	127	138		TM C	Cytosolic
LG:233444.9:200	2JAN18	139	161			cytosolic ransmer

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TABLE 4

		Chart	Ictor	Frame	Domain	Topology
SEQ ID	Template ID	Start	Stop		Type	Topology
NO:		740	194		TM	Extracellular
98	LG:233444.9:2002JAN18	162	217		TM	Transmembrane
98	LG:233444.9:2002JAN18	195	237	-	TM	Cytosolic
98	LG:233444.9:2002JAN18	218		-	TM	Transmembrane
98	LG:233444.9:2002JAN18	238	260	_	TM	Extracellular
98	LG:233444.9:2002JAN18	261	274		TM	Transmembrane
98	LG:233444.9:2002JAN18	275	297		TM	Cytosolic
98	LG:233444.9:2002JAN18	298	430	- 	TM	
98	LG:233444.9:2002JAN18	431	453			Transmembrane Extracellular
98	LG:233444.9:2002JAN18	454	472		TM	Transmembrane
98	LG:233444.9:2002JAN18	473	495		TM	
98	LG:233444.9:2002JAN18	496	671		TM	Cytosolic
98	LG:233444.9:2002JAN18	672	694		TM	Transmembrane
98	LG:233444.9:2002JAN18	695	761		TM_	Extracellular
98	LG:233444.9:2002JAN18	762	784		TM	Transmembrane
98	LG:233444.9:2002JAN18	785	802	ļ	TM	Cytosolic
98	LG:233444.9:2002JAN18	803	825		TM	Transmembrane
98	LG:233444.9:2002JAN18	826	910		TM	Extracellular
98	LG:233444.9:2002JAN18	1	194		TM	Extracellular
98	LG:233444.9:2002JAN18	195	217		TM	Transmembrane
98	LG:233444.9:2002JAN18	218	228		TM	Cytosolic
98	LG:233444.9:2002JAN18	229	251		TM	Transmembrane
98	LG:233444.9:2002JAN18	252	801		TM	Extracellular
98	LG:233444.9:2002JAN18	802	824		TM	Transmembrane
98	LG:233444.9:2002JAN18	825	910		TM	Cytosolic
99	LG:234824.7:2002JAN18	1	. 1193		TM	Extracellular
99	LG:234824.7:2002JAN18	1194	1216		TM	Transmembrane
99	LG:234824.7:2002JAN18	1217	1227		TM	Cytosolic
99	LG:234824.7:2002JAN18	1228	1250		TM	Transmembrane
99	LG:234824.7:2002JAN18	1251	1253		TM	Extracellular
99	LG:234824.7:2002JAN18	1254	1273		TM	Transmembrane
99	LG:234824.7:2002JAN18	1274	1279		TM	Cytosolic
99	LG:234824.7:2002JAN18	1280	1302		TM	Transmembrane
99	LG:234824.7:2002JAN18	1303	1942		TM	Extracellular
99	LG:234824.7:2002JAN18	1943	1965		TM	Transmembrane
99	LG:234824.7:2002JAN18	1966	1984		TM	Cytosolic
99	LG:234824.7:2002JAN18	1	1945_		TM	Extracellular
99	LG:234824.7:2002JAN18	1946	1968		TM	Transmembrane
99	LG:234824.7:2002JAN18	1969	1984		TM	Cytosolic
99	LG:234824.7:2002JAN18	1	1936		TM	Extracellular
99	LG:234824.7:2002JAN18	1937	1956		TM	Transmembrane
99	LG:234824.7:2002JAN18	1957	1984		TM	Cytosolic
99	LG:234824.7:2002JAN18	2337	2408	forward 3	SP	
99	LG:234824.7:2002JAN18	2337	2390	forward 3	SP	
99	LG:234824.7:2002JAN18	2508	2579	forward 3	SP	
99	LG:234824.7:2002JAN18	2721	2822	forward 3	SP	
99	LG:234824.7:2002JAN18	4626	4676	forward 3	SP	
99	LG:234824.7:2002JAN18	2508	2582	forward 3	SP	
100	LG:235708.23:2002JAN18	1	67		TM	Extracellular
100	LG:235708.23:2002JAN18	68	90		TM	Transmembrane

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TABLE 4

			DLE 4			
SEQ ID	Template ID	Start	Stop			Topology
NO:	•		,		Туре	
100	LG:235708.23:2002JAN18	91	101		TM	Cytosolic
100	LG:235708.23:2002JAN18	102	124		TM	Transmembrane
100	LG:235708.23:2002JAN18	125	1270		TM	Extracellular
101	LG:236649.14:2002JAN18	1	457		<u>TM</u>	Extracellular
101	LG:236649.14:2002JAN18	458	480		TM	Transmembrane
101	LG:236649.14:2002JAN18	481	492		TM	Cytosolic
101	LG:236649.14:2002JAN18	493	515		TM	Transmembrane
101	LG:236649.14:2002JAN18	516	529		TM	Extracellular
101	LG:236649.14:2002JAN18	530	552		TM	Transmembrane
101	LG:236649.14:2002JAN18	553	604		TM	Cytosolic
101	LG:236649.14:2002JAN18	1	391		TM	Extracellular
101	LG:236649.14:2002JAN18	392	414		TM	Transmembrane
101	LG:236649.14:2002JAN18	415	455		TM	Cytosolic
101	LG:236649.14:2002JAN18	456	478		TM	Transmembrane
	LG:236649.14:2002JAN18	479	539		TM	Extracellular
101	LG:236649.14:2002JAN18	540	562		TM	Transmembrane
101	LG:236649.14:2002JAN18	563	603		TM	Cytosolic
101	LG:236649.14:2002JAN18	1	392		TM	Cytosolic
101	LG:236649.14:2002JAN18	393	415		TM	Transmembrane
101		416	454		TM	Extracellular
101	LG:236649.14:2002JAN18	455	477		TM	Transmembrane
101	LG:236649.14:2002JAN18	478	603		TM	Cytosolic
101	LG:236649.14:2002JAN18	1	300		TM	Cytosolic
102	LG:332474.7:2002JAN18	301	320		TM	Transmembrane
102	LG:332474.7:2002JAN18	321	321		TM	Extracellular
102	LG:332474.7:2002JAN18		963	forward 1	SP	
102	LG:332474.7:2002JAN18	892	494	forward 3	SP	
102	LG:332474.7:2002JAN18	429	482	forward 3	SP	
102	LG:332474.7:2002JAN18	429	512	forward 3	SP	
102	LG:332474.7:2002JAN18	429		forward 3	SP	
102	LG:332474.7:2002JAN18	429	491	forward 3	SP	
102	LG:332474.7:2002JAN18	429	488	Horwara 3	TM	Extracellular
103	LG:335727.8:2002JAN18	1	346		TM	Transmembrane
103	LG:335727.8:2002JAN18	347	369		TM	Cytosolic
103	LG:335727.8:2002JAN18	370	377			Extracellular
103	LG:335727.8:2002JAN18	1	214		TM	Transmembrane
103	LG:335727.8:2002JAN18	215	237		TM	Cytosolic
103	LG:335727.8:2002JAN18	238	341		TM	Transmembrane
103	LG:335727.8:2002JAN18	342	360		TM	
103	LG:335727.8:2002JAN18	361	377	<u> </u>	TM	Extracellular
103	LG:335727.8:2002JAN18	25	90	forward 1		France of Health
104	LG:481983.1:2002JAN18	11	19		TM	Extracellular
104	LG:481983.1:2002JAN18	20	42		TM	Transmembrane
104	LG:481983.1:2002JAN18	43	450_		TM	Cytosolic
104	LG:481983.1:2002JAN18	451	469		TM	Transmembrane
104	LG:481983.1:2002JAN18	470	488		TM	Extracellular
104		489	511		TM	Transmembrane
104		512	571		TM	Cytosolic
104		572	594		TM	Transmembrane
104		595	603		TM	Extracellular

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TABLE 4

SEQ ID	Template ID	Start	Stop	Frame	Domain	Topology
NO:	i i i i i i i i i i i i i i i i i i i			1	Type	, ,
104	LG:481983.1:2002JAN18	604	626		TM	Transmembrane
104	LG:481983.1:2002JAN18	627	928		TM	Cytosolic
104	LG:481983.1:2002JAN18	929	951		TM	Transmembrane
104	LG:481983.1:2002JAN18	952	1382	1	TM	Extracellular
104	LG:481983.1:2002JAN18	1383	1405		TM	Transmembrane
104	LG:481983.1:2002JAN18	1406	1611		TM	Cytosolic
104	LG:481983.1:2002JAN18	1612	1634		TM	Transmembrane
104	LG:481983.1:2002JAN18	1635	2147		TM	Extracellular
104	LG:481983.1:2002JAN18	2148	2170		TM	Transmembrane
104	LG:481983.1:2002JAN18	2171	2271		TM	Cytosolic
104	LG:481983.1:2002JAN18	1	36		TM	Extracellular
104	LG:481983.1:2002JAN18	37	59		TM	Transmembrane
104	LG:481983.1:2002JAN18	60	159		TM	Cytosolic
104	LG:481983.1:2002JAN18	160	179		TM	Transmembrane
104	LG:481983.1:2002JAN18	180	188		TM	Extracellular
104	LG:481983.1:2002JAN18	189	206		TM	Transmembrane
104	LG:481983.1:2002JAN18	207	243		TM	Cytosolic
104	LG:481983.1:2002JAN18	244	266		TM	Transmembrane
104	LG:481983.1:2002JAN18	267	280		TM	Extracellular
104	LG:481983.1:2002JAN18	281	303		TM	Transmembrane
104	LG:481983.1:2002JAN18	304	447		TM	Cytosolic
104	LG:481983.1:2002JAN18	448	470		TM _	Transmembrane
104	LG:481983.1:2002JAN18	471	927		TM	Extracellular
104	LG:481983.1:2002JAN18	928	950		TM	Transmembrane
104	LG:481983.1:2002JAN18	951	969		TM _	Cytosolic
104	LG:481983.1:2002JAN18	970	992		TM	Transmembrane
104	LG:481983.1:2002JAN18	993	1160		TM	Extracellular
104	LG:481983.1:2002JAN18	1161	1183		TM	Transmembrane
104	LG:481983.1:2002JAN18	1184	1216		TM	Cytosolic
104	LG:481983.1:2002JAN18	1217	1239		TM	Transmembrane
104	LG:481983.1:2002JAN18	1240	2271		TM	Extracellular
104	LG:481983.1:2002JAN18	1465	1524	forward 1	SP	
104	LG:481983.1:2002JAN18	1465	1518	forward 1	SP	
104	LG:481983.1:2002JAN18	1465	1509	forward 1	SP	
104	LG:481983.1:2002JAN18	3116	3160	forward 2	SP	
104	LG:481983.1:2002JAN18	1271	1345	forward 2	SP	
104	LG:481983.1:2002JAN18	1271	1351	forward 2	SP	
104	LG:481983.1:2002JAN18	570	626	forward 3	SP	
104	LG:481983.1:2002JAN18	831	905	forward 3	SP	
104	LG:481983.1:2002JAN18	1833	1892	forward 3	SP	<u> </u>

GEO ID NO:/Template ID	Component Span
1/LG:1447398.9:2002JAN18	1513-1947; 1446-1810; 1232-1808; 1394-1805; 1349-1804; 1330-1905; 137-1905; 137-1905; 137-1905; 137-1905; 1313-1610; 1232-1482; 993-1458; 1159-1447; 1024-1415; 1424-1798; 1540-1780; 1321-1619; 1313-1610; 1232-1482; 993-1458; 106-1161; 798-1161; 909-1161; 174-1389; 998-1286; 601-1232; 836-1212; 724-1168; 905-1160; 662-1100; 627-1100; 693-1100; 914-1161;927-1161; 777-1161; 746-1160; 739-1153; 646-1160; 662-1100; 635-990; 562-907;
	/ 15-1100; 020-1100, 000-1100; 000-1302; 77-416; 701-535; 296-784; 351-630; 502-903; 580-815; 1-624; 194-484
2/LG:201488.3:2002JAN18	1-226; 75-328; 77-245; 176-447; 245-319; 270-319; 270-319; 271-326; 516-857; 659-891; 351-551; 354-936; 474-578; 539-792; 556-905; 610-853; 613-874; 613-864; 616-857; 659-891; 351-551; 354-936; 474-578; 539-720-1281; 734-992; 736-786; 774-1344; 749-899; 760-1023; 817-664-785; 676-1210; 714-967; 720-1281; 734-992; 736-786; 676-1210; 714-967; 720-1281; 734-992; 736-786; 676-1210; 718-967; 720-1281; 734-992; 736-786; 744-1344; 749-899; 760-1023; 817-
	979; 839-1111; 847-1117; 864-1318; 888-1581; 896-1438; 904-1203; 718-1102; 723-1109-1050; 987-1567; 1035-1116; 1059-1527; 1081-1228; 1088-1292; 1085-1476; 1107-1351; 1160-
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	-	LG:1447398.9:2002JAN18	9
	2	LG:201488.3:2002.IAN18	T
	က	LG:288410.6:2002.JAN18	High and Track 23%, N. Cardiovascular System - 12%, Unclassified/Mixed - 11%
	4	LG:7682817,1;2002,1AN18	Cardiovasoular 8 and 1987 - 1984 Endocrine System - 22%
	5	LG:7685059.6:2002.JAN18	Liver - 37% et.i., 23%, Hemic and Immune System - 25%, Exocrine Glands - 25%
	9	LG:7689671.1:2002JAN18	Digestive System 75% No.
	7	LG:7689684.1:2002JAN18	Digestive System 30% Library
	œ	LG:7762669.1:2002JAN18	Pancrage - 82% Diagrafic and Immune System - 30%, Male Genitalia - 20%, Female Genitalia -
	6	LG:965822.1:2002JAN18	Skin - 43%, Spring Organia - 18%
	10	LG:006394,31:2002.JAN18	Septe Orders 156/ 2: 24%, Germ Cells - 19%
	11	LG:018258.1:2002.JAN18	Hinday Track 130, E. L. Control of the Color Cells - 13%
	12	LG:027320.5:2002JAN18	Germ Celfs - 33% Embracial - 29%, Digestive System - 21%
	13		Skin - 13%
			Stomotopopulation 2000 11 1 1 2000
	15		Exocrine Cland: 35% 11-1 22%, Unclassified/Mixed - 13%
		1	Sense Ordans 24% Embrand/Mixed - 25%, Endocrine System - 20%
		7	Inclusified Missal 28% 0 "
		100	Unclassified Military 20%, Cardiovascular System - 31%, Endocrine System - 31%
	19	-	Liver - 35% Mills 1-35%, Hemic and Immune System - 23%, Nervous System - 23%
	20		Germ Cells, 21%, Emb.
	21	or	liver 25% Embracia 21.
		LG:1097492.2:2002.JAN18	Highery Track 100
		G:1099945.26;2002JAN18	LG:1099945,26:2002JAN18 Miscriloskolatal 8
		LG:110016.1:2002JAN18	Miscrifoskolatal System - 13%, Germ Cells - 11%
- 1		α	wideky dietrib, to d
		LG:118836.26:2002.IAN18	Endocrino Quatern 1900 C
		7~	Male Centrals 17% 6 "
*	28 L(G:1347461.28:2002JAN181	LG:1347461,28:2002JAN18 Unclassified Milyad 2007 11.
7		LG:1383494.16:2002JAN18 Sense	Sense Orders 179/ 1902 1100
اري		LG:1400155.1:2002JAN18	
۲	31 [[6	1	Indicate Surface 420/ 11:
		7	Liver - 23% Hodgand M. C. J. 125. Female Gentfalla - 11%, Nervous System - 11%
		7	25.2, directionalised - 17%, Respiratory System - 15%

Tissue Distribution	wide	18 Nervous System - 26%, Skin - 16%, Female GenItalia - 13%	Sense			æ			3 Unclassified/Mixed - 40%, Respiratory System - 20%, Nervous System - 20%		3 Sense Organs - 19%, Nervous System - 19%		8	9 Pancreas - 36%, Respiratory System - 23%, Urinary Tract - 12%, Skin - 12%	80		Г	18 Sense Organs - 24%, Digestive System - 12%, Liver - 10%			Г									Uncl		8 Sense Organs - 21%, Liver - 13%
Template ID	LG:1452619.1:2002JAN18	LG:1453417.6:2002JAN18	LG:148485.8:2002JAN18	LG:1502670.1:2002JAN18	LG:206593,3:2002JAN18	LG:228273.22:2002JAN18	LG:228319.2:2002JAN18	LG:229165.16:2002JAN18	LG:230895,9:2002JAN18	LG:233552.5:2002JAN18	LG:234430.7:2002JAN18	LG:236659.1:2002JAN18	LG:236767.26:2002JAN18	LG:237489.7:2002JAN18	LG:238218.20:2002JAN18	LG:239939.14:2002JAN18	LG:242288.11:2002JAN18	LG:242491.29:2002JAN18	LG:243488.41:2002JAN18	LG:247792.18:2002JAN18	LG:253193,17:2002JAN18	LG:257088.20:2002JAN18	LG:265552.1:2002JAN18	LG:275355.12:2002JAN18	LG:280014.1:2002JAN18	LG:299937.3:2002JAN18	LG:311197.3:2002JAN18	LG:321069.2:2002JAN18	LG:330900.8:2002JAN18	LG:330931.9:2002JAN18	LG:330985.1:2002JAN18	LG:332027.9:2002JAN18
SEQ ID NO: Template ID	33	34	35	36	37	38	39	40	4	42	43	4	45	46	47	48	49	20	51	52	53	54	55	56	57	28	29	9	19	62	63	49

SEO ID NO.	Tomplate ID	Tissue Distribution
.]_	3002 IAN18	Unclassified/Mixed - 24%, Nervous System - 22%, Maie Genitalia - 19%
3 %	.05.20023/1018	Germ Cells - 36%, Male Genitalia - 12%
26	1G:340580.16:2002JAN18	Female Genitalia - 11%
89	LG:350272.6:2002JAN18	Embryonic Structures - 13%, Musculoskeletal System - 12%
69	LG:397228.1:2002JAN18	Unclassified/Mixed - 42%, Cardiovascular System - 33%, Male Gentfalla - 17%
20	LG:401325.41:2002JAN18	Nervous System - 19%, Skin - 12%, Unclassified/Mixed - 10%
71	LG:402029.14:2002JAN18	Respiratory System - 17%, Male Genitalia - 15%, Digestive System - 13%
72	LG:407233.2:2002JAN18	Endocrine System - 28%, Hemic and Immune System - 24%, 5kiii - 21%
73	LG:407346.1:2002JAN18	Sense Organs - 29%, Nervous System - 26%, Hemic and Immune System - 11%
74	LG:407689.7:2002JAN18	Digestive System - 14%, Nervous System - 11%, Hemic and Immune System - 11%
75	LG:407700.1:2002JAN18	Skin - 21%, Digestive System - 10%
76	LG:410461.92:2002JAN18	Musculoskeletal System - 19%, Liver - 19%, Endocrine System - 10%
77	LG:411043.3:2002JAN18	Pancreas - 16%, Exocrine Glands - 12%, Nervous System - 12%
78	LG:438690.47:2002JAN18	widely distributed
79	LG:444677.81:2002JAN18	Sense Organs - 31%, Urinary Tract - 15%, Skin - 12%, Male Genifalia - 12%
8	LG:457464.24:2002JAN18	Skin - 13%, Hemic and Immune System - 10%
81	LG:7684793.15:2002JAN18	
82	LG:7687485.1:2002JAN18	Cardiovascular System - 26%, Musculoskeletal System - 20%, Utinary 11act - 14%
83	LG:7689661.4:2002JAN18	Musculoskeletal System - 28%, Urinary Iract - 11%, Nervous system - 10%, Endocurie system - 20%
84	LG:7690373.1:2002JAN18	Male Genitalia - 40%, Nervous System - 40%, Hemic and Immune System - 20%
85	LG:7696560.1:2002JAN18	Nervous System - 53%, Pancreas - 47%
98	LG:7698190.26:2002JAN18 widely distributed	widely distributed
87	LG:7763560.12:2002JAN18 Sense Organs - 10%	Sense Organs - 10%
88	LG:7763587.20:2002JAN18	LG:7763587.20:2002JAN18 Sense Organs - 23%, Musculoskeletal System - 12%
89	LG:899263.10:2002JAN18	Female Genitalia - 16%, Nervous System - 16%, Embryonic Structures - 14%, Exocili 19 Glatias - 14%, Evol.
		14%
8	LG:977837.31:2002JAN18	Liver - 41%, Respiratory System - 29%, Unclassified/Mixed - 29%
16	LG:978560.13:2002JAN18	Germ Cells - 11%, Embryonic Structures - 11%
92	LG:979390.2:2002JAN18	Pancreas - 25%, Connective fissue - 19%, Liver - 19%
93	LG:983019.1:2002JAN18	Embryonic Structures - 37%, Germ Cells - 29%, Unclassified/Mixed - 11%
94	LG:997202.7:2002JAN18	Skin - 11%, Exocrine Glands - 10%
95	LG:998756.3:2002JAN18	Germ Cells - 20%

SEQ ID NO:	SEQ ID NO: Template ID	Tissue Distribution
%	LG:103460.28:2002JAN18 Germ Cells - 10%	Germ Cells - 10%
76	LG:1501505.19:2002JAN18	LG;1501505.19;2002JAN18 Musculoskeletal System - 50%, Male Genitalia - 33%, Digestive System - 17%
86	LG:233444.9:2002JAN18	Sense Organs - 14%, Urinary Tract - 12%
66	LG:234824.7:2002JAN18	Germ Cells - 21%, Sense Organs - 18%
9	LG:235708.23:2002JAN18	LG:235708.23:2002JAN18 Digestive System - 25%, Liver - 22%, Male Genitalia - 16%
101	LG:236649.14:2002JAN18	LG;236649,14;2002JAN18 Germ Cells - 15%, Musculoskeletal System - 10%
102	LG:332474.7:2002JAN18	Urinary Tract - 56%, Nervous System - 25%, Female Genitalia - 13%
103	1 1	Endocrine System - 19%, Cardiovascular System - 18%, Musculoskeletal System - 18%
104	LG:481983.1:2002JAN18	Germ Cells - 12%

nber Probability Score Annotation	predicted protein washing.	CAMPA14 IIKE	dJ899C14.1 (novel pionell sitting) 1	LIM domains containing profession in the MGD, source key:MGI: 1352502,		evidence:1555~purditive	LIM domains containing protein 1	Similar to amyotrophic lateral scierosis 2 (juvernie) Cirio riosomo 1881.	candidate 7	ALS2CR7	KIAA0834 protein	Unknown (protein for MeC.2/349)	putative putative	QJ0886-8.1 (SITIIII 10 1100801101 10 1100 10 10 10 10 10 10 10 10 10	unnamed protein product	unnamed protein product									Unknown (projein lot Method: concentral translation supplied by			Unknown (piolein for wood of the control of the con			hypothelical profession for MGC:28436)	
Probabilit	1.00E-89	5622 2.00E-51	1527 2.00E-51	1.00E-98	2.00E-98		2.00E-96	6557 1.00E-87	•	1.00E-87	6.00E-51	72170 3.00E-20	1145 2.00E-15	1.00E-08	7.00E-91	4.00E-31	16783 2.00E-23	4.00E-10	58286 5.00E-10	60554 8.00E-10	39498 2.00E-77	73014 2.00E-76	54469 4.00E-76	54722 2.00E-22	, 2.00E-17	2.00E-17	60908 2.00E-40	3 5.00E-18	3 8.00E-10	1.00E-79	7 2.00E-75	5 3.00E-75
GI Number	174	g15485622	g10241527	g6599307 1.00E-98	g12836264 2.00E-98		d6599070	g24416557		g15823642 1.00E-87	157	g20072170	g2231145	g6522712	g21750807 7.00E-91	g10439850 4.00E-31	g18916783	g930123	g21758286	g22760554	g21739498	g23273014	g21754469	g21754722	g15080547 2.00E-17	g1049301	g22760908	g13623633 5.00E-18	g16551840 8.00E-10	g6808105	g21739797 2.00E-79	g193531/5 3.00E-79
Stop G		1039 g	1039 g	674 9	674 g		674	1		1132		1373		1373 (766			079		0/9	604		604	456	456	456	- 1	720	720	2323	2323	2323
			536		3		c	14		614	614	870	870	870	206	206	206	452	452	452	26	26	59	133	133	133	1	-	_	1367	1367	1367
Length Start	168	168	168	224	224	 	700	173	2	173	173	891	168	168	187	187	187	73	73	73	182	182	182	108	108	108	240	240	240	319	319	319
Frame		0	2	3	65	>	-	2 5	٧	0	10	1 60	(n)	3	0	2	10	10	10	2	2	2	2	-	-	-	-	_	-	2	2	2
SFO ID NO:		105	105	106	305	3	70.	2 2) -	107	107	2 2	108	108	3 2	601	001	011	011	110	2 =			112	112	112	113	113	113	114	114	114

Lenath Start Stop GI Number Probability Score	263 2 790 g19483967 1.00E-110	2 790	0.00 O.001	2 790 g3/5//19 Z.UUE-6/	3 1349 912839186 0.00	1349 016563765 0.00	3 1349 g23094020 1.00E-108	1 1803 95689563 0.00	1 1803 g12407441 0.00	g5834582 0.00	'81 6.00E-85	389 4.00E-82	170 17 526 g12833285 3.00E-65 HesB-like domain confamiling professional sources in the second source in the second	13 8.00E-37	456 773 g7022229 8.00E-37	106 456 773 g4894380 8.00E-37	334 95 1096 g14165480 1.00E-178	334 95 1096 g12053195 1.00E-178	334 95 1096 g5305706 1.00E-177	109 11 337 g17381941 2.00E-43	109 11 337 g999454 7.00E-30	109 11 337 g903934 7.00E-30	519 1 1557 g23342615 0.00	1 1557 g21899842 0.00	1 1557 g21886479 0.00	132 3 398 g21336362 3.00E-24	132 3 398 g21757193 5.00E-22	132 3 398 g7023216 4.00E-17	
enath Sta	63 2								501	501	170 17										109	109	519 1	519	519				
Frame	_	2		2			2 2		1	1	2			 3					2			0		-	-	. 60	ر س	3	
SEO ID NO:	115			115	116	114	911	117	117	117	118	118	118	119	119	119	120	120	120	121	121	121	120	122	122	123	123	123	

TABLE 7

SEQ ID NO:		Frame Length Start		Stop	IDN IO	nber Probability Score Annotation	Annotation
124	. 2	524	290	1861	g7019945	1.00E-179	unnamed protein product
124		524	230	1861	g22760462 1.00E-165	1.00E-165	unnamed protein product
125	-	309	685	1611	g3540177	1.00E-109	F23269 2
125		309	685	1611	g5080758	1.00E-108	BC331191 1
125		306	685	1611	g12855931	5931 5.00E-63	data source:SPTR, source key:P16374, evidence:ISS~putative~similar to
		000					ZINC FINGER PROTEIN 60 (ZFP-60) (ZINC FINGER PROTEIN MFG-3)
	7	339	1979	2995	g10047329	7329 1.00E-178	KIAA1626 protein
		339	- 1	2995	g18256873 1.00E-161	1.00E-161	Unknown (protein for IMAGE:4016433)
	2		626	2995	g7022610	1.00E-151	unnamed protein product
127		163	7	490	g12842823 2.00E-38	2.00E-38	data source:SPTR, source key:P02403, evidence:ISS~homolog to 60S
							RIBOSOMAL PROTEIN L37 (G1,16)~putative
	2	163	2	490	g57121	4.00E-38	ribosomal protein L37
			2	490	g461232	4.00E-38	ribosomal protein L37
128		352	259	1314	g16552019	1.00E-109	unnamed profein product
128	_		259	1314	g12844321 1.00E-96	1.00E-96	data source:SPTR, source key:Q9ER30, evidence:ISS-homolog to KFI CH-
							RELATED PROTEIN 1 (KEL-LIKE PROTEIN 23) (SARCOSIN) JUI HAHIVA
			259	1314	g16306813 8.00E-39	8.00E-39	
	2			- 1	g7582193	5.00E-22	dynein light chain 1 protein D.C-1
			2		g470344	5.00E-22	C. elegans DLC-1 protein (corresponding seculance 17445 0)
	2	138			g4103059	5.00E-22	protein inhibitor of nitric oxide synthase
		809	1148	2971	g7959175 0.00	00.0	KIAA1457 protein
	2	809	1148	2971	g6599224 0.00	0.00	hypothetical profein
			1148		g12667438 0.00	00.0	NIR3
131			460	2541	g4929551 0.00	00.0	CGI-40 protein
131	2	694	460	2541	g23274030 0.00	00.0	Similar to CGI-40 protein
					g22761032 0.00	00.0	unnamed protein product
					g23242933 (00.0	Unknown (protein for MGC:46349)
132 3		_	231	1433	g21755437 0.00	00.0	unnamed protein product
				3	g20515159 3	159 3.00E-15	conserved hypothetical protein
					g4929669 2.00E-70	2.00E-70	CGI-100 protein
			T	-	g16741027 2	027 2.00E-70	CGI-100 protein
133	3	141	474	8%	g11596068 2	368 2.00E-70	dJ976O13.1 (CGI-100 protein)

 				Π	- T			T	T			\neg			\top	T	T		T					
Annotation dJ73M23.2 (NAD+-dependent succinic semialdehyde dehydrogenase (SSADH, EC 1.2.1.24))	aldehyde dehydrogenase 5 family, member A1 (succinate-settilater) yac dehydrogenase)	KIAA1473 protein Similar to zinc finger protein 91 (HPF7, HTF10)	hematopoletic cell derived zinc illiger protes.	unnamed protein product	unnamed protein product	Unknown (ProcessMGD, source:MGD, source:MGD, source:CUG triplet repeat, RNA binding protein 1~data source:MGD, source	key:MGI:1342295, evidence:lb>~purding	RNA-binding profess by a contract of the contr	FLAMINGO I	Similar to D.melanogaster cadherin-related tumor supplessor	Unknown (protein for MGC:40579)	unnamed protein product	CG1271-PA	data source;SPIR, source key, r412/3, 3, 3, 3, 3, 1, 2, 3, 1, 2, 3, 1, 3, 3, 1, 3, 3, 1, 3, 3, 1, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3, 3,	RIBOSYLATION FACTOR-EINE 1801	purglive protor-like protein 1	ADF-IIDOsylanov received	Lind in 1961 (Section 1) (Section 1) (Section 1) (Section 2)	unamed profesion product	KIAA0681 protein	dJ138B7.3.2 (lethal (3) mallgnant brain 141101 (10) 113.2	(Drosophila) homolog (Isotorm 2) (Kirkhuda 1)	I(3)mbt protein homolog	
Length Start Stop GI Number Probability Scores 340 392 1411 g4164365 0.00	2 340 392 1411 g3766467 0.00 2 340 392 1411 g21708023 0.00	229 312 998 g7959207 3.00E-38	g3342002 9.00E-36	407 1	1 407 1 1221 g10440210 1.00E-128	529 2 1588 92247	529 2 1588	1588	254 8988	2254 8988 g21929188	2254 8988	14/0 9210	1470	490 1 1470))	100 2 571 9607	100 2 571	153 692 1150	153 692 1150	692 1150	608 583	583 2406	- 500 583 2406 Q3811111 0.00	
SEQ ID NO: Frame 134 2	134	135	135		136	136	137		137		138	139	139	139	140		047	140	141	141	142	142		142

	SEQ ID NO; Frame Length Start	JO; Fra	me Ler	igth Sto	art Stop	<u>0</u>	Number	1ABLE 7 Number Probability Score Apportation	IABLE 7
	143	4	265	5 646				752073 1.00E-148	indialists distance between the second secon
	143	-	265	646	6 1440	Γ	5041142	g16041142 11.00E-133	a indition protein product
	143	-	265				1017813	G14017813 1 ODE-88	MAN 1758
	4	2	687				012024744 0 00	000	TA 222
!	144	2	687	σ.	2068	1	71707	200	DUCID COOL AT
	144	2	687		3	1	G3605004 0.00	0.00	KEI protein
	145	_	478		1434	1	542080F 0.00	0.00	IA p63 alpha
	145	-	478	-	1434		450	0.00	KIAA1918 protein
<u> </u>	145	-	478	-	1434	2 2	143y	0.00	UDP-GallNAc:polypeptide N-acetylgalactosaminyl transferase
	146	-	946	274	1		0.00	0.00	UDP-GallNAc:polypeptide, N-acetylgalactosaminyltransferase
	146	_	946	274		1010	430072 0.00	0.00	KIAA1604 protein
	146	_	946	274	Τ		438214 0.00	00.00	Unnamed protein product
	147	6	318	6	Τ	277	7,00	000	undmed protein product
	:	<u> </u>	2	2		/cob	Ç 77.	1.00E-170	dJ37E16.5 (novel protein similar to nitrophenylphosphatases from various
17	147	6	318	63	056	7010	453107	353107 1 00F 170	organisms)
 79	147	6	318	0 65	3,55	212	70100		hypothetical protein dJ37E16.5
	148	3	1632		300	Τ	314002499 3.00E-99	3.00E-99	Similar to hypothetical protein dJ37E16.5
<u>L_</u>	148	(r.	1635		404	Т	912003890 0.00	00.	polybromo-1
<u>L</u>	148	0 00	1,435	2 0	4070		9120838/5 0.00	00.	polybromo-1
	2 2	2	1007	Т	4878		g16607752 0.00	00:	Unnamed profes product
	149	N	736	7	2209		g12836469 0.00	00.	data source:SPTR, source key:Q9HCJ3, evidence:ISS~homolog to KIAA1579
	149	2	736	2	2200	70000	00 0 001000	00	PROJEIN (FRAGMENT)-putative
	149	2	736	2	2200		766810 0.00	30	Unknown (protein for IMAGE:5113697)
	350	-	323	157	1105	1	00017	00:	Unknown (protein for MGC:46327)
	155	-	323	157	1125		4155/ 1.00E-16	.00E-161	anion fransporter/exchanger-9
L_	150	-	303	157	1105	7	915341550 1.00E-161	.UUE-161	putative anion transporter
_	15.	-	270	<u>2</u> ,	27	\neg	g14/8/223 2,00E-29		prestin
	151	-	070	- -	1884		9287 0.00		KIAA1513 protein
<u>L</u>	151	-	809	- -	1887	9201	g2U146520 1.00E-101		SLIP003
	152	60	183	32/	270	0140/	70188 3.00E-27		Hypothetical protein W02H5,4
_	152	6	183	327	272	3000	921730740 3,00E-92		unnamed protein product
	152	6.	183	32.7	0770	9200/	7324U 3.UUE-92		similar to putative
]		2	3	1024	7/0	1015/	g15/95887 3.00E-92		unnamed protein product

umber Probability Score Annotation	KIAA1142 protein	PAK4 protein	serine/threonine kinase	unnamed profein product	unnamed profein product	P26 protein	Similar to HTPAP protein	НТРАР	data source:SPTR, source key:Q9VND5, evidence:ISS~putative~related to	Hoknown (protein for MGC:37640)	unnamed profein product	Unknown (protein for MGC:33943)	E74-like factor 1 (ets domain transcription factor)	Ets-family transcription factor ELF1	transcription factor Elf-1	euchromatic histone methyltransferase 1	unnamed protein product	KIAA1876 protein	RIKEN cDNA 2010002A20 gene	Immunoglobulin domain containing protein~data source:Pfam, source Immunoglobulin domain containing protein~data source:Pfam, source	Immunoglobulin domain containing protein~data source:Pfam, source	key:PF00047, evidence:ISS~putative	ATP-binding cassette sub-family A member 9	ATP-binding cassette sub-family A member 9	ATP-binding cassette A9	Unknown (protein for MGC:23949)	claudin 14~data source:MGD, source key:MGI:1860425,	claudin 14	unnamed protein product
Probability Sco	1,00E-132	1.00E-132	1.00E-132	075 1.00E-103	3.00E-99	682 5.00E-92	541 1.00E-119	757 3.00E-85	3,00E-58	00.0	0.00	0.00	00.00	0.00	0.00	0.00	0.00	0.00	1.00E-56	1.00E-56	961 1.00E-50		450 1.00E-122	407 1.00E-122	1.00E-121	1.00E-80	1.00E-30	2508 1.00E-30	2414 1.00E-155
GI Number	de32662	g4164385	g4101587	g22760075	g15209772 3.00E-99	g20377682	g21542541	g13182757	g12844263 3.00E-58	003073603	g10041649 0.00	g23271317 0.00	g20988140 0.00	g15010800 0.00	g11995007 0.00	g20372683 0.00	g10434623 0.00	g20522002 0.00	g18490501 1.00E-56	g12843712 1.00E-56	g12841961		g23451450	g23451407	g17223624 1.00E-121	g18088580	g12860781 1.00E-30	g13452508	g16549414
Stop		1512	1512	262			953		953	1858	1858	1858	2309	2309	2309	3601	3601	3601	622	622	623	1	849	849	849	1005	1005	1005	1112
	1	499	499	25			3	က	3	a	0 00	8	1197	1197	1197	686	686	686	152	152	152	!	92	2	10	559	559	559	270
Length	338	338	338	191	161	191	317	317	317	217	617	617	371	371	371	871	871	871	157	157	157	<u> </u>	280	280	280	149	149	149	281
Frame		_		_			3		က	c	2 6	2	3	8	8	2	2	2	2	2	0	1	-	_	_	_			က
SEQ ID NO: Frame Length Start	153	153	153	154	154	154	155	155	155		156				157		158				150	<u> </u>	160	160	160	161	161	161	162

SEQ ID NO: Frame Length Start Stop GI Number Probability Score Annotation	1112 g13938	338 1.00E-77	703 312 2420 g13811938 0.00 dJ1056H1.2.1 (novel protein similar to mitogen inducible protein MIG-2	703 312 2420 g23273527 0.00 Unknown (protein for MGC;46404)	g16878257 0.00	g24059753 5.00E-64	128 439 822 g12856090 9.00E-62 data source:SPTR, source key:Q9Z2B2, evidence:ISS~putative~similar to	e key:6	BRAIN MITOCHONDRIAL CARRIER PROTEIN-1 (BMCP-1)	3751 g21756799 0.00 unnamed protein product		3751 g6650822 0.00	705 1 2115 g13365895 0.00 hypothetical protein		705 1 2115 g1842216 8.00E-17 Zfp64		15 0.00	g2967654 0.00	389 132 1298 g12832845 1.00E-161 Domain of unknown function DUF36 containing protein~data	1298 g217490	389 132 1298 g21410962 1.00E-70 Unknown (protein for MGC;32708)	381 83 1225 g5114351 1.00E-165 RING finger protein terf	1,00E-165	83 1225 g5114353 1.00E-118	3 1979 g14017797 0.00	659 3 1979 g13529311 1.00E-126 Similar to hypothetical protein FLJ23119	659 3 1979 g10439701 1.00E-109 unnamed protein product	219 1253 1909 g8096557 1.00E-84 PBX1B	
hStart	270	270	312	312	312	439	439	439		2	2	2	1	1	1	3	3	3	132	132	132	83	83	83	3	3	3	1253	
) Lengtl	281	281	703	703	703	128	128	 128		1250	1250	1250	705	705	705	630	630	630	386	389	386	381	381	381	629	626	626	219	
Frame	3	3	က	 3	3	1	_	1		2	2	2	1	1		3	3	3	ည	က	3	2	2	2	3	က	က	2	
SEQ ID NO	162	162	163	163	163	164	164	164		165	165	165	166	166	166	167	167	167	168	168	168	169	169	169	170	170	170	171	

ber Probability Score Annotation	1,00E-84 homeobox protein		366 1.00E-124 Zinc finger, C3HC4 type (RING finger) containing protein-data source; Pfam, source key:PF00097, evidence: ISS-putative	235 1.00E-82 Similar to RIKEN cDNA 0610037N03 gene			339 2.00E-13 zinc finger protein 147 (estrogen-responsive finger protein)			77 1.00E-143 BAP2-beta protein			300 2.00E-99 Unknown (protein for IMAGE:4984604)			8 0.00 SH2 domain-containing phosphatase anchor protein 2a	8 0.00 enaptin	9 0.00 nesprin-1		5 0.00 hypothetical protein	0.00 KIAA0931 protein	3 0.00 KIAA0606 protein	272 1.00E-139 Unknown (protein for MGC:5352)	731 1,00E-123 data source;SPTR, source key;O46084, evidence;ISS~putative~related to	EG:63B12.4 PROTEIN	0 1.00E-63 Hypothetical protein R07G3.5	169 1,00E-170 unnamed protein product	445 1,00E-167 g1-related zinc finger protein	1.00E-167 g1-related zinc finger protein	8 0.00 unnamed protein product	2 n nn
GI Number Proba	g456109 1.00E-	g22859174 0.00	g12845866 1.00E-	g13477235 1.00E-	g15741221 1.00E-17	g458726 2.00E-	g16877339 2.00E-	g4239984 1.00E-	g4239982 1.00E-143	g4126477 1.00E-	g3327220 1.00E-	g19550878 1.00E-103	g20072000 2.00E-	g16033591 0.00	g18092655 0.00	g16033588 0.00	g22597198 0.00	g24417709 0.00	g22597200 0.00	g13365845 0.00	94589506 0.00	920521103 0.00	g14198272 1.00E-	g12848731 1.00E-		g23820820 1.00E-63	g13185169 1.00E-	g17390445 1.00E-	g6175860 1.00E-167	g21755898 0.00	010000000000000000000000000000000000000
Stop G	ĺ	1315 g		1315 g	318 g	318 g	318 g	1072 g	1072 g	1072 g	1343 g	1343 g			1664 g	1664 g	2816 g	2816 g	2816 g	2804 g	2804 g	2804 g		913 g		913 g	1166 g	1166 g		1291 g	1001
1.	I	2	2	2	_	-	_	2	2	2	546	546	546	255	255	255	3	က	ဗ	21	21	21	2	2		2	202	207	207	218	מנט
Lengt	219	438	438	438	305	106	305	357	357	357	266	266	266	470	470	470	938	938	938	928	928	928	304	304		304	320	320	320	358	250
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TABLE 7

SEQ ID NO: Frame Length Start	: Frame	Length	Start	Stop	GI Number	ber Probability Score Annotation	Annotation
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202	2	247	527	1267	g7023136	36 1.00E-134	unnamed profein product
202	2	247	527	1267	g13278516 1.00E-101		Similar to hypothetical protein FLJ10846
203	2	749	1034	3280	g4239895 0.00		MASL1
203	2	749	1034	3280	g15559745 0.00		Similar to MFH-amplified sequences with leucine-rich tandem repeats 1
203	2	749	1034	3280	g19916243 3.00E-14	3.00E-14	hypothetical protein
204		330	910	1899	g21756391 9.00E-27	9.00E-27	unnamed protein product
205	- 1	301	133	1035	g20067239 1.00E-118	1.00E-118	putative regulation protein GS3
205	1	301	133	1035	g12859941 3.00E-56	3.00E-56	evidence:NAS~hypothetical protein~putative
205	1	301	133	1035	g5052516 7.00E-25	7.00E-25	BcDNA.GH03108
206	1	213	1	636	g21739500 1.00E-08		hypothetical protein
206	-	213	-	636	g456269	3.00E-05	zlnc finger protein 30
206		213	_	639	g12851705	705 3.00E-05	data source:MGD, source key:MGI:99178, evidence:ISS~putative~zinc
							finger protein 30
207	2	322	2	296	g15559324 1.00E-180	1.00E-180	Unknown (protein for IMAGE:4309224)
207	2	322	2	296	g16551666 1	666 1.00E-173	unnamed protein product
207	2	322	2	296	g21739611 1.00E-109	1.00E-109	hypothetical protein
208	3	238	381	1094	g21410798 1.00E-80	1.00E-80	Unknown (protein for IMAGE:4396549)
208	3	238	381	1094	g19526448 2.00E-69		TRH4
208	3	238	381	1094	g21618502 4.00E-68		Similar to RIKEN cDNA 2310081H14 gene

Program	Description	Reference	Parameter Threshold
ABI FACTURA	A program that removes vector sequences and masks ambiguous bases in nucleic acid sequences.	Applied Biosystems, Foster City, CA.	
ABI/PARACEL FDF	A Fast Data Finder useful in comparing and annotating amino acid or nucleic acid sequences.	Applied Biosystems, Foster City, CA; Paracel Inc., Pasadena, CA.	Mismatch <50%
ABI AutoAssembler	A program that assembles nucleic acid sequences.	Applied Biosystems, Foster City, CA.	
BLAST	A Basic Local Alignment Search Tool useful in sequence similarity search for amino acid and nucleic acid sequences. BLAST includes five functions: blastp, blastn, blastx, tblastn, and tblastx.	Fool useful in sequence Altschul, S.F. et al. (1990) J. Mol. Biol. 215:403-ESTs: Probability value= 1.0E-8 or less; and nucleic acid 410; Altschul, S.F. et al. (1997) Nucleic Acids Full Length sequences: Probability value= 1.0E-10 or less	ESTs: Probability value= 1.0E-8 or less; Full Length sequences: Probability value= 1.0E-10 or less
FASTA	A Pearson and Lipman algorithm that searches for similarity between a query sequence and a group of sequences of the same type. FASTA comprises as least five functions: fasta, fasta, fastx, and ssearch.	Pearson, W.R. and D.J. Lipman (1988) Proc. Natl. Acad Sci. USA 85:2444-2448; Pearson, W.R. (1990) Methods Enzymol. 183:63-98; and Smith, T.F. and M.S. Waterman (1981) Adv. Appl. Math. 2:482-489. ESTs: fasta E value=1.06E-6; Assembled ESTs: fasta E value=6.76E-6; Assembled ESTs: fasta E value=6.76E-6; Assembled ESTs: fasta E value=6.76E-6; Assembled ESTs: fasta E value=1.06E-6; Assembled ESTs: fasta Identity=95% or greater and Nath. 2:482-489.	ESTs: fasta E value=1.06E-6; Assembled ESTs: fasta Identity= 95% or greater and Match length=200 bases or greater; fastx E value=1.0E-8 or less; Full Length sequences: fastx score=100 or greater
BLIMPS	A BLocks IMProved Searcher that matches a sequence against those in BLOCKS, PRINTS, DOMO, PRODOM, Acids Res. 19:6565-6572; Henikoff, J.G. and S. and PFAM databases to search for gene families, sequence homology, and structural fingerprint regions. Henikoff, S. and J.G. Henikoff, 1.G. and S. and S. and S. Henikoff, J.G. and S. Henikoff, J.G. and S. and S. Henikoff, J.G. and S. Henikoff, J.G. and S. Henikoff, J.G. and S. and S. Henikoff, J.G. and S. Henikoff, J.G. and S. Acids Res. 19:6565-6572; Henikoff, J.G. and S. Henikoff, J.G. and J.G. an		Probability value= 1.0E-3 or less
HMMER	An algorithm for searching a query sequence against hidden Markov model (HMM)-based databases of protein family consensus sequences, such as PFAM.	Krogh, A. et al. (1994) J. Mol. Biol. 235:1501- PFAM hits: Probability value= 1.0E-3 of 1531; Sonnhammer, E.L.L. et al. (1988) Nucleic less; Acids Res. 26:320-322; Durbin, R. et al. (1998) Signal peptide hits: Score= 0 or greater Our World View, in a Nutshell, Cambridge Univ. Press, pp. 1-350.	PFAM hits: Probability value= 1.0E-3 or less; Signal peptide hits: Score= 0 or greater

Program	Description	Reference	Parameter Threshold
ProfileScan	An algorithm that searches for structural and sequence motifs in protein sequences that match sequence patterns Gribskov, M. et al. (1988) CABIOS 4:61-66; motifs in protein sequences that match sequence patterns Gribskov, M. et al. (1989) Methods Enzymol. 183:146-159; Bairoch, A. et al. (1997) Nucleic Acids Res. 25:217-221.	Gribskov, M. et al. (1988) CABIOS 4:61-66; Gribskov, M. et al. (1989) Methods Enzymol. 183:146-159; Bairoch, A. et al. (1997) Nucleic Acids Res. 25:217-221.	Normalized quality score SGCG-specified "HIGH" value for that particular Prosite motif. Generally, score=1.4-2.1.
Phred	A base-calling algorithm that examines automated sequencer traces with high sensitivity and probability.	Ewing, B. et al. (1998) Genome Res. 8:175-185; Ewing, B. and P. Green (1998) Genome Res. 8:186-194.	·
Phrap	A Phils Revised Assembly Program including SWAT and CrossMatch, programs based on efficient implementation of the Smith-Waterman algorithm, useful in searching sequence homology and assembling DNA sequences.	Smith, T.F. and M.S. Waterman (1981) Adv. Appl. Math. 2:482-489; Smith, T.F. and M.S. Waterman (1981) J. Mol. Biol. 147:195-197; and Green, P., University of Washington, Seattle, WA.	Score= 120 or greater; Match length= 56 or greater
Consed	A graphical tool for viewing and editing Phrap assemblies.	Gordon, D. et al. (1998) Genome Res. 8:195- 202.	
SPScan	A weight matrix analysis program that scans protein sequences for the presence of secretory signal peptides.	Nielson, H. et al. (1997) Protein Engineering 10:1-6; Claverie, J.M. and S. Audic (1997) CABIOS 12:431-439.	Score=3.5 or greater
ТМАР	A program that uses weight matrices to delineate transmembrane segments on protein sequences and determine orientation.	Persson, B. and P. Argos (1994) J. Mol. Biol. 237:182-192; Persson, B. and P. Argos (1996) Protein Sci. 5:363-371.	
TMHMMER	A program that uses a hidden Markov model (HMM) to delineate transmembrane segments on protein sequences and determine orientation.	Sonuhammer, E.L. et al. (1998) Proc. Sixth Intl. Conf. On Intelligent Systems for Mol. Biol., Glasgow et al., eds., The Am. Assoc. for Artificial Intelligence (AAAI) Press, Menlo Park, CA, and MIT Press, Cambridge, MA, pp. 175-182.	

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TABLE
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Program	Description	Reference	Parameter Threshold
Motifs	A program that searches amino acid sequences for patterns that matched those defined in Prosite.	Bairoch, A. et al. (1997) Nucleic Acids Res. 25:217-221; Wisconsin Package Program Manual, version 9, page M51-59, Genetics Computer Group, Madison, WI.	

CLAIMS

What is claimed is:

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- An isolated polynucleotide comprising a polynucleotide sequence selected from the group
 consisting of:
 - a) a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104,
 - b) a naturally occurring polynucleotide sequence having at least 90% sequence identity to a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104,
 - c) a polynucleotide sequence complementary to a),
 - d) a polynucleotide sequence complementary to b), and
 - e) an RNA equivalent of a) through d).
 - 2. An isolated polynucleotide of claim 1, comprising a polynucleotide sequence selected from the group consisting of SEQ ID NO:1-104.

3. An isolated polynucleotide comprising at least 60 contiguous nucleotides of a polynucleotide of claim 1.

- 4. A composition for the detection of expression of disease detection and treatment molecule polynucleotides comprising at least one of the polynucleotides of claim 1 and a detectable label.
 - 5. A method for detecting a target polynucleotide in a sample, said target polynucleotide comprising a sequence of a polynucleotide of claim 1, the method comprising:
 - a) amplifying said target polynucleotide or fragment thereof using polymerase chain reaction amplification, and
 - b) detecting the presence or absence of said amplified target polynucleotide or fragment thereof, and, optionally, if present, the amount thereof.
 - 6. A method for detecting a target polynucleotide in a sample, said target polynucleotide comprising a sequence of a polynucleotide of claim 1, the method comprising:
 - a) hybridizing the sample with a probe comprising at least 20 contiguous nucleotides comprising a sequence complementary to said target polynucleotide in the sample, and which probe specifically hybridizes to said target polynucleotide, under conditions whereby a hybridization complex is formed between said probe and said target polynucleotide or fragments thereof, and

b) detecting the presence or absence of said hybridization complex, and, optionally, if present, the amount thereof.

- 7. A method of claim 5, wherein the probe comprises at least 30 contiguous nucleotides.
- 8. A method of claim 5, wherein the probe comprises at least 60 contiguous nucleotides.
- 9. A recombinant polynucleotide comprising a promoter sequence operably linked to a polynucleotide of claim 1.
 - 10. A cell transformed with a recombinant polynucleotide of claim 9.
 - 11. A transgenic organism comprising a recombinant polynucleotide of claim 9.
- 12. A method for producing a disease detection and treatment molecule polypeptide, the method comprising:
- a) culturing a cell under conditions suitable for expression of the disease detection and treatment molecule polypeptide, wherein said cell is transformed with a recombinant polynucleotide of claim 9, and
 - b) recovering the disease detection and treatment molecule polypeptide so expressed.
- 13. A purified disease detection and treatment molecule polypeptide (MDDT) encoded by at least one of the polynucleotides of claim 2.
- 14. An isolated antibody which specifically binds to a disease detection and treatment molecule polypeptide of claim 13.
 - 15. A method of identifying a test compound which specifically binds to the disease detection and treatment molecule polypeptide of claim 13, the method comprising the steps of:
 - a) providing a test compound;
 - b) combining the disease detection and treatment molecule polypeptide with the test compound for a sufficient time and under suitable conditions for binding; and
 - c) detecting binding of the disease detection and treatment molecule polypeptide to the test compound, thereby identifying the test compound which specifically binds the disease detection and treatment molecule polypeptide.

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16. A microarray wherein at least one element of the microarray is a polynucleotide of claim3.

- 17. A method for generating a transcript image of a sample which contains polynucleotides, the method comprising the steps of:
 - a) labeling the polynucleotides of the sample,
 - b) contacting the elements of the microarray of claim 16 with the labeled polynucleotides of the sample under conditions suitable for the formation of a hybridization complex, and
 - c) quantifying the expression of the polynucleotides in the sample.

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- 18. A method for screening a compound for effectiveness in altering expression of a target polynucleotide, wherein said target polynucleotide comprises a polynucleotide sequence of claim 1, the method comprising:
- a) exposing a sample comprising the target polynucleotide to a compound, under conditions suitable for the expression of the target polynucleotide,
 - b) detecting altered expression of the target polynucleotide, and
 - c) comparing the expression of the target polynucleotide in the presence of varying amounts of the compound and in the absence of the compound.
 - 19. A method for assessing toxicity of a test compound, said method comprising:
 - a) treating a biological sample containing nucleic acids with the test compound;
 - b) hybridizing the nucleic acids of the treated biological sample with a probe comprising at least 20 contiguous nucleotides of a polynucleotide of claim 1 under conditions whereby a specific hybridization complex is formed between said probe and a target polynucleotide in the biological sample, said target polynucleotide comprising a polynucleotide sequence of a polynucleotide of claim 1 or fragment thereof;
 - c) quantifying the amount of hybridization complex; and
 - d) comparing the amount of hybridization complex in the treated biological sample with the amount of hybridization complex in an untreated biological sample, wherein a difference in the amount of hybridization complex in the treated biological sample is indicative of toxicity of the test compound.
 - 20. An array comprising different nucleotide molecules affixed in distinct physical locations on a solid substrate, wherein at least one of said nucleotide molecules comprises a first

oligonucleotide or polynucleotide sequence specifically hybridizable with at least 30 contiguous nucleotides of a target polynucleotide, said target polynucleotide having a sequence of claim 1.

- 21. An array of claim 20, wherein said first oligonucleotide or polynucleotide sequence is completely complementary to at least 30 contiguous nucleotides of said target polynucleotide.
 - 22. An array of claim 20, wherein said first oligonucleotide or polynucleotide sequence is completely complementary to at least 60 contiguous nucleotides of said target polynucleotide
 - 23. An array of claim 20, which is a microarray.
 - 24. An array of claim 20, further comprising said target polynucleotide hybridized to said first oligonucleotide or polynucleotide.
 - 25. An array of claim 20, wherein a linker joins at least one of said nucleotide molecules to said solid substrate.
 - 26. An array of claim 20, wherein each distinct physical location on the substrate contains multiple nucleotide molecules having the same sequence, and each distinct physical location on the substrate contains nucleotide molecules having a sequence which differs from the sequence of nucleotide molecules at another physical location on the substrate.
 - 27. An isolated polypeptide comprising an amino acid sequence selected from the group consisting of:
 - a) an amino acid sequence selected from the group consisting of SEQ ID NO:105-208,
 - b) a naturally occurring amino acid sequence at least 90% identical to an amino acid sequence selected from the group consisting of SEQ ID NO:105-208,
 - c) a biologically active fragment of an amino acid sequence selected from the group consisting of SEQ ID NO:105-208, and
- d) an immunogenic fragment of an amino acid sequence selected from the group consisting of SEQ ID NO:105-208.
 - 28. An isolated polypeptide of claim 27, comprising a polypeptide sequence selected from the group consisting of SEQ ID NO:105-208.

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<110> INCYTE GENOMICS, INC.; JONES, Anissa L.;
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86/218

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Thr Arg Thr Pro Ala Thr Asp Cys Leu Met Tyr Leu Gln Gly Pro
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Arg Lys Leu Met Thr Gln Gly Gly Tyr Asp Met Val Gln Lys Leu
                 35
                                     40
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Phe Leu Asp Phe Phe Arg Arg Leu Ser Gln Arg Pro Thr Ala
                 50
                                     55
Glu Glu Leu Glu Gln Arg Asn Ile Leu Lys Pro Arg Asn Glu Gln
                 65
                                     70
Glu Glu Glu Glu Lys Arg Glu Ile Lys Arg Arg Leu Thr Arg
                 80
                                     85
                                                          90
Lys Leu Ser Gln Arg Pro Thr Val Glu Glu Leu Arg Glu Arg Lys
                 95
                                    100
                                                         105
Ile Leu Ile Arg Phe Ser Asp Tyr Val Glu Val Ala Asp Ala Gln
                110
                                    115
Asp Tyr Asp Arg Arg Ala Asp Lys Pro Trp Thr Arg Leu Thr Ala
                125
                                    130
                                                         135
Ala Asp Lys Ala Ala Ile Arg Lys Glu Leu Asn Glu Phe Lys Ser
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Thr Glu Met Glu Val His Glu Leu Ser Arg His Leu Thr Arg Phe
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His Arg Pro
<210> 106
<211> 224
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Arg Glu Leu Glu Arg Ala Leu Glu Ala Arg Thr Ala Arg Asp Tyr
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Phe Gly Ile Cys Ile Lys Cys Gly Leu Gly Ile Tyr Gly Ala Gln
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Gln Ala Cys Gln Ala Met Gly Ser Leu Tyr His Thr Asp Cys Phe
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Thr Cys Asp Ser Cys Gly Arg Arg Leu Arg Gly Lys Ala Phe Tyr
                  50
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Asn Val Gly Glu Lys Val Tyr Cys Gln Glu Asp Phe Leu Tyr Ser
                  65
Gly Phe Gln Gln Thr Ala Asp Lys Cys Ser Val Cys Gly His Leu
                  80
Ile Met Glu Met Ile Leu Gln Ala Leu Gly Lys Ser Tyr His Pro
                  95
                                     100
Gly Cys Phe Arg Cys Ser Val Cys Asn Glu Cys Leu Asp Gly Val
                110
                                     115
Pro Phe Thr Val Asp Val Glu Asn Asn Ile Tyr Cys Val Arg Asp
                125
                                     130
Tyr His Thr Val Phe Ala Pro Lys Cys Ala Ser Cys Ala Arg Pro
                140
                                     145
Ile Leu Pro Ala Gln Gly Cys Glu Thr Thr Ile Arg Val Val Ser
                155
                                     160
Met Asp Arg Asp Tyr His Val Ala Cys Tyr His Cys Glu Asp Cys
                170
                                     175
Gly Leu Gln Leu Ser Gly Glu Glu Gly Arg Arg Cys Tyr Pro Leu
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                                     190
Ala Gly His Leu Leu Cys Arg Arg Cys His Leu Arg Arg Leu Gln
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Pro Gly Pro Leu Pro Ser Pro Thr Val His Val Thr Glu Leu
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Leu Ile Cys Gly Leu Ala Arg Ala Lys Ser Ile Pro Ser Gln Thr
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Tyr Ser Ser Glu Val Val Thr Leu Trp Tyr Arg Pro Pro Asp Ala
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Leu Leu Gly Ala Thr Glu Tyr Ser Ser Glu Leu Asp Ile Trp Gly
                                      55
                                                          60
Ala Gly Cys Ile Phe Ile Glu Met Phe Gln Gly Gln Pro Leu Phe
Pro Gly Val Ser Asn Ile Leu Glu Gln Leu Glu Lys Ile Trp Glu
                 80
                                      85
Val Leu Gly Val Pro Thr Glu Asp Thr Trp Pro Gly Val Ser Lys
                 95
                                     100
Leu Pro Asn Tyr Asn Pro Glu Trp Phe Pro Leu Pro Thr Pro Arg
                110
                                     115
Ser Leu His Val Val Trp Asn Arg Leu Gly Arg Val Pro Glu Ala
                125
                                     130
Glu Asp Leu Ala Ser Gln Met Leu Lys Gly Phe Pro Arg Asp Arg
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145

Val Ser Ala Gln Glu Ala Leu Val His Asp Tyr Phe Ser Ala Leu

140

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155
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Pro Ser Gln Leu Tyr Gln Thr Ser
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Leu Gly Ala Leu Ser Arg Met Arg Phe Glu Asp Tyr Ala Asn Ala
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Glu Ala His Val His Gln Pro Ala His Gln Val Gln Gly Ile Cys
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Cys Leu Gln Gly Leu Glu Arg Ser Arg Leu Val Trp Val Leu Ser
                 35
                                      40
Val Pro Arg Arg Cys Arg Pro Arg Gly His His Ser Gly Gln Ala
                 50
                                      55
Phe His Cys Ser Arg Ala Gln Arg Leu Leu Gly Glu Gln Asp Gln
                 65
                                      70
Gln Ala Pro His His Pro Leu Gln Gly Asp Arg Pro Leu Gln Leu
                 80
                                      85
Cys Ala Cys Ala Pro His Pro Cys Ala Gln Gly His Trp Cys His
                . 95
                                     100
Leu Gly Pro Arg Tyr Ser Leu Glu Cys Cys Cys Leu Val Ala Gly
                110
                                     115
Ile Asp Asp Cys Tyr Thr Ser Ala Arg Ser Cys Thr Ala Thr Leu
                125
                                     130
Gly Asn Phe Ala Lys Thr Thr Phe Asp Ala Ile Ser Ile Asp Leu
                140
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Gln Leu Pro Asp Pro Arg Pro Leu Glu Glu Asp Cys Val His Gln
Val Ser Leu
<210> 109
<211> 187
<212> PRT
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Gly Pro Leu Ser Pro Gly Pro Tyr Gln Cys Arg Pro Ser Leu Pro
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Ala Gln Leu Tyr Pro Gln Ser Leu Met Ala Ala Ala Thr Leu Arg
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Thr Pro Thr Gln Gly Thr Val Thr Phe Glu Asp Val Ala Val His
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Phe Ser Trp Glu Glu Trp Gly Leu Leu Asp Glu Ala Gln Arg Cys
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Leu Tyr Arg Asp Val Met Leu Glu Asn Leu Ala Leu Leu Thr Ser
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                                      70
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Leu Asp Val His His Gln Lys Gln His Leu Gly Glu Lys His Phe
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Arg Ser Asn Val Gly Arg Ala Leu Phe Val Lys Thr Cys Thr Phe
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His Val Ser Gly Glu Pro Ser Thr Cys Arg Glu Val Gly Lys Asp
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                                      115
 Phe Leu Ala Lys Leu Gly Phe Leu His Gln Gln Ala Ala His Thr
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 Gly Glu Gln Ser Asn Ser Lys Ser Asp Val Gly Ala Ile Ser His
                 140
                                      145
                                                          150
 Arg Gly Lys Thr His Cys Asn Cys Gly Glu His Thr Lys Ala Phe
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                                      160
 Ser Gly Lys His Thr Leu Val Gln Gln Gln Arg Thr Leu Thr Thr
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 Glu Arg Cys Tyr Ile Arg Ser
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Arg Glu Ser Glu Gly Lys Glu Asp Gly Gln Cys Glu Glu Ile Phe
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Ser Leu Val Pro Asn Gly Ile Val Lys Thr Thr Phe Thr Gly Val
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Lys Ser Cys Glu Ser Ser Val Cys Glu Glu Gly Asn Met Asp His
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Ser Ser Leu Asn Cys Cys Ile Arg Ala Asp Thr Gly His Lys Ser
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Asp Glu Cys Gln Gln His Arg Ser His Ile Ser Ser Val
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Pro Lys Gln Gly Ile Arg Val Trp Ser Pro Arg His Pro Glu Asn
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Phe Leu Gly Ile Glu Ser Arg Pro Pro Val Leu Ser Leu Ser Pro
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Ile Leu Leu Tyr Thr Cys Glu Met Phe Gln Asp Pro Val Ala Phe
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Asp Asp Val Ala Val Asn Phe Thr Gln Glu Glu Trp Ala Leu Leu
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Asp Ile Ser Gln Arg Lys Leu Tyr Lys Glu Val Met Leu Glu Thr
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Phe Arg Asn Leu Thr Ser Val Gly Lys Ser Trp Lys Asp Gln Asn
                 80
                                      85
Ile Glu Tyr Glu Tyr Gln Asn Pro Arg Arg Asn Phe Arg Ser Leu
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                                                         105
Ile Glu Lys Lys Val Asn Glu Ile Lys Asp Asp Ser His Cys Gly
                110
                                     115
Glu Thr Phe Thr Gln Val Pro Asp Asp Arg Leu Asn Phe Gln Glu
                125
                                    130
Lys Lys Ala Ser Pro Glu Ile Lys Ser Cys Asp Ser Phe Val Cys
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145
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                140
Gly Glu Val Gly Leu Gly Asn Ser Ser Phe Asn Met Asn Ile Arg
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Gly Asp Ile Gly His Lys Ala Tyr Glu Tyr Gln Glu Tyr Gly Pro
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Lys Pro
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<213> Homo sapiens
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Val Ala Phe Glu Asp Val Ala Val Asn Phe Thr His Glu Glu Trp
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Ala Leu Leu Gly Pro Ser Gln Lys Asn Leu Tyr Arg Asp Val Met
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                                                           45
                 .35
Leu Glu Asn Phe Gln Asn Leu Ala Ser Leu Gly Tyr Pro Leu His
                                      55
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Thr Pro His Leu Ile Ser Gln Trp Glu Gln Glu Glu Asp Leu Gln
                 65
Thr Val Lys Arg Glu Leu Ile Gln Gly Ile Phe Met Gly Glu His
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Arg Glu Gly Lys Asn Pro Trp Glu Lys Leu Phe Trp Leu Gly Glu
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Lys Ile Asn
<210> 113
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Pro Arg Ser Ser Arg Arg Val Trp Ala Ala Tyr Thr Glu Gly Lys
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Lys Lys Pro Ser Phe Leu Gly Lys Cys Arg Lys Pro Ala Ser Gly
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Arg Ser Leu Arg Ser Pro Gln Gly Gly Ala Leu Ala Ala Gln Arg
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                  35
                                       40
Ala Arg Phe Pro Ala Gly Glu Pro Arg Asn Gly Gly Ala Gly Gly
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                  50
                                       55
Gly Asp Ser Glu Asp Pro Arg Leu Gly Phe Pro Thr Trp Ile Arg
                  65
                                       70
Ser Ala Trp Gly Phe Asp Pro His Pro Gly Ala Ala Pro Arg Arg
                  80
                                       85
Ser Trp Ala Ala Arg Ala Phe Gly Leu Arg His Arg Gln Arg His
                  95
                                      100
Leu Glu Ala Gly Ala Ser Gly Arg Leu Cys Leu Thr Cys Leu Leu
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                                      115
Glu Gly Asn Thr Gly Lys Pro Gly Leu Ala Val Thr Leu Val Thr
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Asn Met Ser Gln Asp Ser Val Thr Phe Ala Asp Val Ala Val Asn
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Phe Thr Lys Glu Glu Trp Thr Leu Leu Asp Pro Ala Gln Arg Asn
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Leu Tyr Arg Asp Val Met Leu Glu Asn Ser Arg Asn Leu Ala Phe
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                                     175
Ile Asp Trp Ala Thr Pro Cys Lys Thr Lys Asp Ala Thr Pro Gln
                 185
                                     190
Pro Asp Ile Leu Pro Lys Arg Thr Phe Pro Glu Ala Asn Arg Val
                 200 -
Cys Leu Thr Ser Ile Arg Phe Pro Ala Leu His Ile Lys Arg Arg
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Leu Glu Met Pro Gln Asn Arg Gly Thr Thr Gln Ala Gly Gly Glu
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Gly Ser Ala Phe Arg Val Pro Cys Pro Ile Leu Glu Gly Pro Ala
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Ala Gly Ser Arg Pro Arg Leu Ser Glu Ala Met Gly Ile Gln Ser
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Ala Glu Leu Pro Pro Glu Glu Ser Asp Ser Ser Arg Val Asp Phe
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Gly Ser Ser Glu Arg Leu Gly Ser Trp Gln Glu Lys Glu Glu Asp
                 65
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Ala Arg Pro Asn Ala Ala Ala Pro Ala Leu Gly Pro Val Gly Leu
                 80
                                      85
Glu Ser Asp Leu Ser Lys Val Arg His Lys Leu Arg Lys Phe Leu
                 95
                                     100
Gln Arg Arg Pro Thr Leu Gln Ser Leu Arg Glu Lys Gly Tyr Ile
                110
                                     115
Lys Asp Gln Val Phe Gly Cys Ala Leu Ala Ala Leu Cys Glu Arg
                125
                                     130
Glu Arg Ser Arg Val Pro Arg Phe Val Gln Gln Cys Ile Arg Ala
                140
                                     145
Val Glu Ala Arg Gly Leu Asp Ile Asp Gly Leu Tyr Arg Ile Ser
                155
                                     160
Gly Asn Leu Ala Thr Ile Gln Lys Leu Arg Tyr Lys Val Asp His
                170
                                     175
Asp Glu Arg Leu Asp Leu Asp Asp Gly Arg Trp Glu Asp Val His
                185
                                     190
Val Ile Thr Gly Ala Leu Lys Leu Phe Phe Arg Glu Leu Pro Glu
                200
                                     205
Pro Leu Phe Pro Phe Ser His Phe Arg Gln Phe Ile Ala Ala Ile
                215
                                     220
Lys Leu Gln Asp Gln Ala Arg Arg Ser Arg Cys Val Arg Asp Leu
                230
                                     235
Val Arg Ser Leu Pro Ala Pro Asn His Asp Thr Leu Arg Met Leu
                245
                                     250
                                                         255
Phe Gln His Leu Cys Arg Val Ile Glu His Gly Glu Gln Asn Arg
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                                     265
                                                         270
Met Ser Val Gln Ser Val Ala Ile Val Phe Gly Pro Thr Leu Leu
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280
Arg Pro Glu Val Glu Glu Thr Ser Met Pro Met Thr Met Val Phe
Gln Asn Gln Val Val Glu Leu Ile Leu Gln Gln Cys Ala Asp Ile
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Phe Pro Pro His
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 Leu Val Ala Ile Leu Cys Val Leu Val Val Trp Ile Phe Lys Asn
 Ala Asp Arg Ser Met Glu Lys Lys Lys Gly Glu Pro Arg Thr Arg
 Ala Glu Ala Arg Pro Trp Val Asp Glu Asp Leu Lys Asp Ser Ser
 Asp Leu His Gln Ala Glu Glu Asp Ala Asp Glu Trp Gln Glu Ser
  Glu Glu Asn Val Glu His Ile Pro Phe Ser His Asn His Tyr Pro
  Glu Lys Glu Met Val Lys Arg Ser Gln Glu Phe Tyr Glu Leu Leu
  Asn Lys Arg Arg Ser Val Arg Phe Ile Ser Asn Glu Gln Val Pro
  Met Glu Val Ile Asp Asn Val Ile Arg Thr Ala Gly Thr Ala Pro
  Ser Gly Ala His Thr Glu Pro Trp Thr Phe Val Val Lys Asp
  Pro Asp Val Lys His Lys Ile Arg Lys Ile Ile Glu Glu Glu Glu
   Glu Ile Asn Tyr Met Lys Arg Met Gly His Arg Trp Val Thr Asp
   Leu Lys Lys Leu Arg Thr Asn Trp Ile Lys Glu Tyr Leu Asp Thr
   Ala Pro Ile Leu Ile Leu Ile Phe Lys Gln Val His Gly Phe Ala
   Ala Asn Gly Lys Lys Val His Tyr Tyr Asn Glu Ile Ser Val
   Ser Ile Ala Cys Gly His Pro Ala Ser Cys Pro Ala Glu Cys Ser
   Leu Val Thr Val Thr Asn Asn Pro Leu Asn Val Ala Ser Asp Glu
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   Gly Cys Pro Trp Ala Ala Arg Thr
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   Gly Glu Ala Gly Arg Ala Pro Asp Ser Asp Gly Gly Ser Asp Ala
  Asp Ser Glu Val Gly Pro Gly Ser Pro Thr Arg Thr Ala Glu Ala
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  Ala Glu Glu Met Ala Gly Pro Asn Gln Leu Cys Ile Arg Arg
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  Trp Thr Thr Lys His Val Ala Val Trp Leu Lys Asp Glu Gly Phe
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  Phe Glu Tyr Val Asp Ile Leu Cys Asn Lys His Arg Leu Asp Gly
                                        70
  Ile Thr Leu Leu Thr Leu Thr Glu Tyr Asp Leu Arg Ser Pro Pro
                                        85
  Leu Glu Ile Lys Val Leu Gly Asp Ile Lys Arg Leu Met Leu Ser
                   95
                                       100
  Val Arg Lys Leu Gln Lys Ile His Ile Asp Val Leu Glu Glu Met
                  110
  Gly Tyr Asn Ser Asp Ser Pro Met Gly Ser Met Thr Pro Phe Ile
                                       115
                  125
                                       130
  Ser Ala Leu Gln Ser Thr Asp Trp Leu Cys Asn Gly Glu Leu Ser
                  140
                                       145
  His Asp Cys Asp Gly Pro Ile Thr Asp Leu Asn Ser Asp Gln Tyr
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 Gln Tyr Met Asn Gly Lys Asn Lys His Ser Val Arg Arg Leu Asp
                                      160
                  170
 Pro Glu Tyr Trp Lys Thr Ile Leu Ser Cys Ile Tyr Val Phe Ile
                                      175
                                      190
 Val Phe Gly Phe Thr Ser Phe Ile Met Val Ile Val His Glu Arg
                                      205
 Val Pro Asp Met Gln Thr Tyr Pro Pro Leu Pro Asp Ile Phe Leu
                  215
                                      220
 Asp Ser Val Pro Arg Ile Pro Trp Ala Phe Ala Met Thr Glu Val
                  230
                                      235
 Cys Gly Met Ile Leu Cys Tyr Ile Trp Leu Leu Val Leu Leu Leu
                 245
                                      250
 His Lys His Arg Ser Ile Leu Leu Arg Arg Leu Cys Ser Leu Met
                 260
 Gly Thr Val Phe Leu Leu Arg Cys Phe Thr Met Phe Val Thr Ser
                                      265
                 275
 Leu Ser Val Pro Gly Gln His Leu Gln Cys Thr Gly Lys Ile Tyr
                                     280
                                     295
 Gly Ser Val Trp Glu Lys Leu His Arg Ala Phe Ala Ile Trp Ser
Gly Phe Gly Met Thr Leu Thr Gly Val His Thr Cys Gly Asp Tyr
                                     325
Met Phe Ser Gly His Thr Val Val Leu Thr Met Leu Asn Phe Phe
                 335
                                     340
Val Thr Glu Tyr Thr Pro Arg Ser Trp Asn Phe Leu His Thr Leu
Ser Trp Val Leu Asn Leu Phe Gly Ile Phe Phe Ile Leu Ala Ala
                                     355
                 365
                                     370
His Glu His Tyr Ser Ile Asp Val Phe Ile Ala Phe Tyr Ile Thr
                380
Thr Arg Leu Phe Leu Tyr Tyr His Thr Leu Ala Asn Thr Arg Ala
                                     385
                                     400
Tyr Gln Gln Ser Arg Arg Ala Arg Ile Trp Phe Pro Met Phe Ser
                410
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Phe Phe Glu Cys Asn Val Asn Gly Thr Val Pro Asn Glu Tyr Cys
Trp Pro Phe Ser Lys Pro Ala Ile Met Lys Arg Leu Ile Gly
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<210> 117

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Pro Pro Arg Leu Ile Ser Val Gln Thr Met Gln Arg Gly Asn Met
Asn Cys Gly Ala Phe Gln Ala His Gln Met Arg Leu Ala Gln Asn
Ala Ala Arg Ile Pro Gly Ile Pro Arg His Ser Gly Pro Gln Tyr
 Ser Met Met Gln Pro His Leu Gln Arg Gln His Ser Asn Pro Gly
 His Ala Gly Pro Phe Pro Val Val Ser Val His Asn Thr Thr Ile
 Asn Pro Thr Ser Pro Thr Thr Ala Thr Met Ala Asn Ala Asn Arg
 Gly Pro Thr Ser Pro Ser Val Thr Ala Ile Glu Leu Ile Pro Ser
 Val Thr Asn Pro Glu Asn Leu Pro Ser Leu Pro Asp Ile Pro Pro
 Ile Gln Ala Asn Val Val Pro Met Met His Ser Trp Tyr Glu Phe
 Gly Ala Arg Glu Lys Thr Gln Asp Gln Asn Val Leu Glu Asp Ala
  Gly Ser Ser Ser Leu Asp Asn Leu Leu Ser Arg Tyr Ile Ser Gly
  Ser His Leu Pro Pro Gln Pro Thr Ser Thr Met Asn Pro Ser Pro
                                      190
  Gly Pro Ser Ala Leu Ser Pro Gly Ser Ser Gly Leu Ser Asn Ser
  His Thr Pro Val Arg Pro Pro Ser Thr Ser Ser Thr Gly Ser Arg
                                       220
  Gly Ser Cys Gly Ser Ser Gly Arg Thr Ala Glu Lys Thr Ser Leu
  Ser Phe Lys Ser Asp Gln Val Lys Val Lys Gln Glu Pro Gly Thr
                                       250
   Glu Asp Glu Ile Cys Ser Phe Ser Gly Gly Val Lys Gln Glu Lys
   Thr Glu Asp Gly Arg Arg Ser Ala Cys Met Leu Ser Ser Pro Glu
   Ser Ser Leu Thr Pro Pro Leu Ser Thr Asn Leu His Leu Glu Ser
   Glu Leu Asp Ala Leu Ala Ser Leu Glu Asn His Val Lys Ile Glu
                                        310
   Pro Ala Asp Met Asn Glu Ser Cys Lys Gln Ser Gly Leu Ser Ser
                   305
   Leu Val Asn Gly Lys Ser Pro Ile Arg Ser Leu Met His Arg Ser
                    320
                                        340
   Ala Arg Ile Gly Gly Asp Gly Asn Asn Lys Asp Asp Pro Asn
                    335
    Glu Asp Trp Cys Ala Val Cys Gln Asn Gly Gly Asp Leu Leu Cys
                    350
                                         370
    Cys Glu Lys Cys Pro Lys Val Phe His Leu Thr Cys His Val Pro
    Thr Leu Leu Ser Phe Pro Ser Gly Asp Trp Ile Cys Thr Phe Cys
                    395
```

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Arg Asp Ile Gly Lys Pro Glu Val Glu Tyr Asp Cys Asp Asn Leu
   Gln His Ser Lys Lys Gly Lys Thr Ala Gln Gly Leu Ser Pro Val
   Asp Gln Arg Lys Cys Glu Arg Leu Leu Tyr Leu Tyr Cys His
                                       430
   Glu Leu Ser Ile Glu Phe Gln Glu Pro Val Pro Ala Ser Ile Pro
                                       445
  Asn Tyr Tyr Lys Ile Ile Lys Lys Pro Met Asp Leu Ser Thr Val
                                       460
                   470
  Lys Lys Lys Leu Gln Lys Lys His Ser Gln His Tyr Gln Ile Pro
                                       475
  Asp Asp Phe Val Ala Asp Val Arg Leu Ile Phe Lys Asn Cys Glu
                                       490
                                      505
  Arg Phe Asn Glu Met Met Lys Val Val Gln Val Tyr Ala Asp Thr
                  515
  Gln Glu Ile Asn Leu Lys Ala Asp Ser Glu Val Ala Gln Ala Gly
                                      520
  Lys Ala Val Ala Leu Tyr Phe Glu Asp Lys Leu Thr Glu Ile Tyr
                                      535
  Ser Asp Arg Thr Phe Ala Pro Leu Pro Glu Phe Glu Gln Glu Glu
                                      550
  Asp Asp Gly Glu Val Thr Glu Asp Ser Asp Glu Asp Phe Ile Gln
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                                      580
  Pro Arg Arg Lys Arg Leu Lys Ser Asp Glu Arg Pro Val His Ile
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 Lys
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 Phe Leu His Tyr His Pro Pro Pro Thr Gln Gly Trp Trp Trp Arg
 Lys Met Ala Ala Ala Trp Gly Ser Ser Leu Thr Ala Ala Thr Gln
Arg Ala Val Thr Pro Trp Pro Arg Gly Arg Leu Leu Thr Ala Ser
Leu Gly Pro Gln Ala Arg Arg Glu Ala Ser Ser Ser Pro Glu
Ala Gly Glu Gly Gln Ile Arg Leu Thr Asp Ser Cys Val Gln Arg
                                     55
Leu Leu Glu Ile Thr Glu Gly Ser Glu Phe Leu Arg Leu Gln Val
                                     85
Glu Gly Gly Cys Ser Gly Phe Gln Tyr Lys Phe Ser Leu Asp
Thr Val Ile Asn Pro Asp Asp Arg Val Phe Glu Gln Gly Gly Ala
                                    100
Arg Val Val Asp Ser Asp Ser Leu Ala Phe Val Lys Gly Ala
                                    115
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160

165

Gln Val Asp Phe Ser Gln Glu Leu Ile Arg Ser Ser Phe Gln Val Leu Asn Asn Pro Gln Ala Gln Gln Gly Cys Ser Cys Gly Ser Ser

Phe Ser Ile Lys Leu

170

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Val Cys Arg Asn Ser Tyr Phe Tyr Leu Pro Ser Leu Ser Glu Ser
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Lys His Cys Leu Arg Ile Gln His Thr Phe Cys Phe Leu Thr Cys
              · 20
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Val Gln Ala Tyr Val His Lys Ser Val Met Glu Glu Leu Lys Arg
                 35
                                      40
Ile Ile Asp Asp Ser Glu Ile Thr Lys Glu Asp Asp Ala Leu Trp
                 50
                                      55
Pro Pro Pro Asp Arg Val Gly Arg Gln Glu Leu Glu Ile Val Ile
                                      70
                 65
Gly Asp Glu His Ile Ser Phe Thr Thr Ser Lys Ile Gly Ser Leu
                 80
                                      85
Ile Asp Val Asn Gln Ser Lys Asp Pro Glu Gly Leu Arg Val Phe
                                     100
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Tyr
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Val Tyr Phe Ala Ala Pro Ser Ala Phe Glu Lys Met Ser Val Thr
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Tyr Asp Asp Ser Val Gly Val Glu Val Ser Ser Asp Ser Phe Trp
                                       25
                                                           30
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Glu Val Gly Asn Tyr Lys Arg Thr Val Lys Arg Ile Asp Asp Gly
                  35
                                       40
                                                           45
His Arg Leu Cys Ser Asp Leu Met Asn Cys Leu His Glu Arg Ala
                  50
                                       55
                                                           60
Arg Ile Glu Lys Ala Tyr Ala Gln Gln Leu Thr Glu Trp Ala Arg
                                       70
                                                           75
                  65
Arg Trp Arg Gln Leu Val Glu Lys Gly Pro Gln Tyr Gly Thr Val
                  80
                                       85
Glu Lys Ala Trp Met Ala Phe Met Ser Glu Ala Glu Arg Val Ser
                  95
                                      100
Glu Leu His Leu Glu Val Lys Ala Ser Leu Met Asn Asp Asp Phe
                                      115
                                                          120
                 110
Glu Lys Ile Lys Asn Trp Gln Lys Glu Ala Phe His Lys Gln Met
                                      130
                 125
Met Gly Gly Phe Lys Glu Thr Lys Glu Ala Glu Asp Gly Phe Arg
                 140
                                      145
Lys Ala Gln Lys Pro Trp Ala Lys Lys Leu Lys Glu Val Glu Ala
                 155
                                      160
Ala Lys Lys Ala His His Ala Ala Cys Lys Glu Glu Lys Leu Ala
                                      175
                                                          180
                 170
Ile Ser Arg Glu Ala Asn Ser Lys Ala Asp Pro Ser Leu Asn Pro
                                                          195
                 185
                                      190
```

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Glu Gln Leu Lys Lys Leu Gln Asp Lys Ile Glu Lys Cys Lys Gln
                200
                                    205
Asp Val Leu Lys Thr Lys Glu Lys Tyr Glu Lys Ser Leu Lys Glu
                215
                                    220
                                                         225
Leu Asp Gln Gly Thr Pro Gln Tyr Met Glu Asn Met Glu Gln Val
                                    235
                230
Phe Glu Gln Cys Gln Gln Phe Glu Glu Lys Arg Leu Arg Phe Phe
                                    250
                245
Arg Glu Val Leu Leu Glu Val Gln Lys His Leu Asp Leu Ser Asn
                260
                                     265
Val Ala Gly Tyr Lys Ala Ile Tyr His Asp Leu Glu Gln Ser Ile
                275
                                     280
Arg Ala Ala Asp Ala Val Glu Asp Leu Arg Trp Phe Arg Ala Asn
                                    295
                290
His Gly Pro Gly Met Ala Met Asn Trp Pro Gln Phe Glu Val Arg
                305
                                     310
Gly Gly Cys Ala His Glu Leu Val Ser Leu Glu Glu Asp Leu Gly
                320
                                     325
Pro Gln Ser Cys
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Gln Trp Lys Lys Pro Val Gln Ile Leu Met Val Gly Ser Cys Lys
Val Thr Ser Val Met Met Leu Leu Gln Arg Leu Met Trp Glu Lys
                 20
Asp Phe Ile Ala Phe Lys Ser Ser Thr Pro His Asn Val Ser Trp
                 35
                                      40
Arg His Glu Thr Asn Gly Ser Val Phe Ile Ser Gln Ile Ile Tyr
                 50
                                      55
Tyr Phe Arg Glu Tyr Ser Trp Ser His His Leu Glu Glu Ile Phe
                 65
                                      70
                                                          75
Gln Lys Val Gln His Ser Phe Glu Thr Pro Asn Ile Leu Thr Gln
                 80
                                      85
Leu Pro Thr Ile Glu Arg Leu Ser Met Thr Arg Tyr Phe Tyr Leu
                 95
                                     100
Phe Pro Gly Asn
<210> 122
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Arg Ala Ala Val Leu Gly Arg Val Arg Gly Gly Leu Ala Ala Glu
                  5
                                      10
Ala Pro Arg Arg Gly Ala Asn Gly Ala Asn Ala Arg Ser Pro
                                      25
                 20
Pro Ala Arg Arg Cys Ala Gly Gly Trp Trp Arg Gly Pro Arg Pro
```

				35					40					45
Thr	Leu	Arg	Thr	Met 50	Thr	Cys	Trp	Leu	Cys 55	Val	Leu	Ser	Leu	Pro 60
Leu	Leu	Leu	Leu	Pro 65	Ala	Ala	Pro	Pro	Pro 70	Ala	Gly	Gly	Cys	Pro 75
Ala	Arg	Суѕ	Glü	Cys 80	Thr	Val	Gln	Thr	Arg 85	Ala	Val	Ala	Суѕ	Thr 90
Arg	Arg	Arg	Leu	Thr 95	Ala	Val	Pro	Asp	Gly 100	Ile	Pro	Ala	Glu	Thr 105
Arg	Leu	Leu	Glu	Leu 110	Ser	Arg	Asn	Arg	Ile 115	Arg	Суѕ	Leu	Asn	Pro 120
Gly	Asp	Leu	Ala	Ala 125	Leu	Pro	Ala	Leu	Glu 130	Glu	Leu	Asp	Leu	Ser 135
Glu	Asn	Ala	Ile	Ala 140	His	Val	Glu	Pro	Gly 145	Ala	Phe	Ala	Asn	Leu 150
Pro	Arg	Leu	Arg	Val 155	Leu	Arg	Leu	Arg	Gly 160	Asn	Gln	Leu	Lys	Leu 165
			_	170			Arg		175					180
_				185	_		Val		190					195
				200			Arg		205					210
				215			Ala		220					225
				230			Cys		235					240
				245			Ser		250					255
				260			Glu		265					270
				275			Ile	•	280					285
				290			Leu		295					300
				305			Ile		310					315
				320			Thr		325					330
				Val 335			Gly		340					345
				350			Gly		355					360
				365			Arg		370					375
				380					385					Ser 390
				395					400					Ala 405
				410					415					Leu 420
				425					430					Val 435
				440					445					Glu 450
				455					460					Gln 465
				470					475					Arg 480
				485					490					Gln 495
HIS	Arg	PLO	val	500		rnr	ser	ALâ	505		ATG	Arg	vai	Leu 510

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Pro Gly Gly Thr Leu Glu Ile Gln Asp
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 <211> 132
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 Gly Asp Pro Pro Arg Pro Gly Phe Cys Pro Ala Arg Ala Asp Ser
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 Arg Lys Ser Gly Ser Gly Ser Arg Gly Val Thr Val Thr Pro Arg
                   20
                                       25
                                                            30
 Arg Ile Asn Ser Gln Arg Leu Met Leu His Glu Lys Ala Thr Lys
                  35
                                       40
                                                            45
 Lys Thr Lys Glu Lys Glu Thr Arg Met Ala Leu Pro Gln Gly Cys
                   50
                                       55
                                                            60
 Leu Thr Phe Lys Asp Val Ala Ile Glu Phe Ser Leu Glu Glu Trp
                  65
                                       70
 Lys Cys Leu Asn Pro Ala Gln Arg Ala Leu Tyr Arg Ala Val Met
                  80
                                       85
 Leu Glu Asn Tyr Arg Asn Leu Glu Ser Val Gly Leu Thr Ser Lys
                  95
                                      100
                                                          105
 Asp Ser Trp Tyr Met Arg Lys Lys Pro Gly Arg Gly Arg Gly Lys
                 110
                                      115
                                                          120
 Gln Arg Arg Gln Glu Trp Phe Phe Leu Arg Val Tyr
                 125
                                      130
 <210> 124
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Pro Cys Thr Lys Arg Asn Gly Asp Cys Leu Tyr Pro Pro Arg Phe
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Ile Ser Trp Pro Glu Val Ile Leu Ala Ser Arg Lys Gly Cys Thr
                  20
                                      25
Ser Ser His His Gln Leu Gln Arg Met Ala Ala Ile Tyr Leu Ser
                  35
                                      40
Arg Gly Phe Phe Ser Arg Glu Pro Ile Cys Pro Phe Glu Glu Lys
                  50
                                      55
                                                           60
Thr Lys Val Glu Arg Met Val Glu Asp Tyr Leu Ala Ser Gly Tyr
                  65
                                      70
Gln Asp Ser Val Thr Phe Asp Asp Val Ala Val Asp Phe Thr Pro
                  80
                                      85
                                                           90
Glu Glu Trp Ala Leu Leu Asp Thr Thr Glu Lys Tyr Leu Tyr Arg
                 95
                                     100
Asp Val Met Leu Glu Asn Tyr Met Asn Leu Ala Ser Val Glu Trp
                110
                                     115
Glu Ile Gln Pro Arg Thr Lys Arg Ser Ser Leu Gln Gln Gly Phe
                125
                                     130
                                                          135
Leu Lys Asn Gln Ile Phe Ser Gly Ile Gln Met Thr Arg Gly Tyr
                140
                                     145
                                                          150
Ser Gly Trp Lys Leu Cys Asp Cys Lys Asn Cys Gly Glu Val Phe
```

Arg Glu Gln Phe Cys Leu Lys Thr His Met Arg Val Gln Asn Gly

```
170
                                    175
Gly Asn Thr Ser Glu Gly Asn Cys Tyr Gly Lys Asp Thr Leu Ser
                185
                                     190
Val His Lys Glu Ala Ser Thr Gly Gln Glu Leu Ser Lys Phe Asn
                                     205
                                                         210
Pro Cys Gly Lys Val Phe Thr Leu Thr Pro Gly Leu Ala Val His
                215
                                     220
Leu Glu Val Leu Asn Ala Arg Gln Pro Tyr Lys Cys Lys Glu Cys
                                     235
                230
Gly Lys Gly Phe Lys Tyr Phe Ala Ser Leu Asp Asn His Met Gly
                                     250
                                                         255
                245
Ile His Thr Asp Glu Lys Leu Cys Glu Phe Gln Glu Tyr Gly Arg
                260
                                     265
                                                          270
Ala Val Thr Ala Ser Ser His Leu Lys Gln Cys Val Ala Val His
                                     280
                275
Thr Gly Lys Lys Ser Lys Lys Thr Lys Lys Cys Gly Lys Ser Phe
                                                         300
                290
                                     295
Thr Asn Phe Ser Gln Leu Tyr Ala Pro Val Lys Thr His Lys Gly
                305
                                     310
                                                          315
Glu Lys Ser Phe Glu Cys Lys Glu Cys Gly Arg Ser Phe Arg Asn
                                     325
                                                          330
                320
Ser Ser Cys Leu Asn Asp His Ile Gln Ile His Thr Gly Ile Lys
                                                          345
                                     340
                335
Pro His Lys Cys Thr Tyr Cys Gly Lys Ala Phe Thr Arg Ser Thr
                                     355
                350
Gln Leu Thr Glu His Val Arg Thr His Thr Gly Ile Lys Pro Tyr
                                                          375
                                     370
                365
Glu Cys Lys Glu Cys Gly Gln Ala Phe Ala Gln Tyr Ser Gly Leu
                                     385
                                                          390
                380
Ser Ile His Ile Arg Ser His Ser Gly Lys Lys Pro Tyr Gln Cys
                                     400
                                                          405
                395
Lys Glu Cys Gly Lys Ala Phe Thr Thr Ser Thr Ser Leu Ile Gln
                                     415
                                                          420
                 410
His Thr Arg Ile His Thr Gly Glu Lys Pro Tyr Glu Cys Val Glu
                                     430
                                                          435
                 425
Cys Gly Lys Thr Phe Ile Thr Ser Ser Arg Arg Ser Lys His Leu
                 440
                                     445
                                                          450
Lys Thr His Ser Gly Glu Lys Pro Phe Val Cys Lys Ile Cys Gly
                                                          465
                                     460
                 455
Lys Ala Phe Leu Tyr Ser Ser Arg Leu Asn Val His Leu Arg Thr
                 470
                                     475
His Thr Gly Glu Lys Pro Phe Val Cys Lys Glu Cys Gly Lys Ala
                 485
                                     490
Phe Ala Val Ser Ser Arg Leu Ser Arg His Glu Arg Ile His Thr
                                     505
                 500
Gly Glu Lys Pro Tyr Glu Cys Lys Asp Met Ser Val Thr Ile
                 515
                                     520
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Asn Ser Tyr Leu Leu Leu Arg His Ser Phe Asn Cys Gly Leu Phe
                                      10
Val Val Val Val Phe Gln Gly Arg Ser Pro Arg Lys Ile Asp Gln
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20
                                       25
                                                           30
 Phe Cys Asn Ser Ser Asn Met Val His Gly Ser Val Thr Phe Arg
                                       40
 Asp Val Ala Ile Asp Phe Ser Gln Glu Glu Trp Glu Cys Leu Gln
                  50
                                       55
 Pro Asp Gln Arg Thr Leu Tyr Arg Asp Val Met Leu Glu Asn Tyr
                  65
                                       70
 Ser His Leu Ile Ser Leu Gly Ser Ser Ile Ser Lys Pro Asp Val
                                       85
 Ile Thr Leu Leu Glu Gln Glu Lys Glu Pro Trp Met Val Val Arg
                  95
                                      100
 Lys Glu Thr Ser Arg Arg Tyr Pro Asp Leu Glu Leu Lys Tyr Gly
                 110
                                                          120
 Pro Glu Lys Val Ser Pro Glu Asn Asp Thr Ser Glu Val Asn Leu
                 125
                                      130
 Pro Lys Gln Val Ile Lys Gln Ile Ser Thr Thr Leu Gly Ile Glu
                 140
                                      145
 Ala Phe Tyr Phe Arg Asn Asp Ser Glu Tyr Arg Gln Phe Glu Gly
                 155
                                      160
 Leu Gln Gly Tyr Gln Glu Gly Asn Ile Asn Gln Lys Met Ile Ser
                 170
                                      175
 Tyr Glu Lys Leu Pro Thr His Thr Pro His Ala Ser Leu Ile Cys
                 185
                                      190
                                                          195
 Asn Thr His Lys Pro Tyr Glu Cys Lys Glu Cys Gly Lys Tyr Phe
                 200
                                      205
 Ser Arg Ser Ala Asn Leu Ile Gln His Gln Ser Ile His Thr Gly
                 215
                                      220
                                                          225
Glu Lys Pro Phe Glu Cys Lys Glu Cys Gly Lys Ala Phe Arg Leu
                 230
                                      235
His Ile Gln Phe Thr Arg His Gln Lys Phe His Thr Gly Glu Lys
                 245
                                      250
Pro Leu Asn Val Thr Asn Val Glu Arg Pro Leu Val Phe Leu Pro
                 260
                                     265
                                                          270
Cys Leu Ile Ala Ile Arg Thr Phe Thr Gln Val Arg Asn Cys Leu
                 275
                                     280
Asn Val Arg Asn Val Gly Ser Pro Leu Ile Val Ala Gln Thr Leu
                 290
                                     295
Phe Asn Ile Arg Val Phe Ile Leu Val
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Ala Arg Phe Pro Gly Ser Thr Gly Tyr Ile Trp Pro Lys Ser Asp
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Ser Leu Gly Ala Leu Val His Ser Pro Val Asn Cys Pro Leu Leu
                 20
                                      25
                                                          30
Gly Phe Ser Ala Val Ser Thr Ser Leu Pro Gln Gly Tyr Leu Trp
                 35
                                      40
Val Gly Gly Gln Glu Gly Ala Gly Gly Gln Val Glu Ile Phe
                 50
Ser Leu Asn Arg Pro Ser Pro Arg Thr Val Lys Ser Phe Pro Leu
                 65
                                      70
                                                          75
Ala Ala Pro Val Leu Cys Met Glu Tyr Ile Pro Glu Leu Glu Glu
                 80
                                      85
Glu Ala Glu Ser Arg Asp Glu Ser Pro Thr Val Ala Asp Pro Ser
```

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100
Ala Thr Val His Pro Thr Ile Cys Leu Gly Leu Gln Asp Gly Ser
Ile Leu Leu Tyr Ser Ser Val Asp Thr Gly Thr Gln Cys Leu Val
Ser Cys Arg Ser Pro Gly Leu Gln Pro Val Leu Cys Leu Arg His
Ser Pro Phe His Leu Leu Ala Gly Leu Gln Asp Gly Thr Leu Ala
Ala Tyr Pro Arg Thr Ser Gly Gly Val Leu Trp Asp Leu Glu Ser
Pro Pro Val Cys Leu Thr Val Gly Pro Gly Pro Val Arg Thr Leu
 Leu Ser Leu Glu Asp Ala Val Trp Ala Ser Cys Gly Pro Arg Val
 Thr Val Leu Glu Ala Thr Thr Leu Gln Pro Gln Gln Ser Phe Glu
 Ala His Gln Asp Glu Ala Val Ser Val Thr His Met Val Lys Ala
 Gly Ser Gly Val Trp Met Ala Phe Ser Ser Gly Thr Ser Ile Arg
 Leu Phe His Thr Glu Thr Leu Glu His Leu Gln Glu Ile Asn Ile
 Ala Thr Arg Thr Thr Phe Leu Leu Pro Gly Gln Lys His Leu Cys
  Val Thr Ser Leu Leu Ile Cys Gln Gly Leu Leu Trp Val Gly Thr
  Asp Gln Gly Val Ile Val Leu Leu Pro Val Pro Arg Leu Glu Gly
  Ile Pro Lys Ile Thr Gln Trp Ala Leu Trp Ala Cys Gly Leu Pro
                  320
  Gly Cys Gly Tyr Gln His Pro Gly Pro
                  335
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   Thr Cys Trp Leu Val Thr Ser Lys Asp Thr Pro Ile Cys Leu Ala
    Pro Trp Leu Ala Ala Pro Thr His Pro Cys Pro Pro Ala
    Gln Ala Pro Asp Pro Lys Glu Glu Val Glu Gly Ala Cys Pro Pro
    Leu Ser Ala Met Gln His Leu Leu Glu Ala Ala Gln Ser Leu Leu
    Thr Ser Val Pro His Leu Ser His Arg Met Gln Lys Met Thr Ser
    Lys Ala Tyr His Leu Gln Lys Ser Thr Cys Gly Lys Cys Gly Tyr
     Pro Ala Lys Arg Lys Arg Lys Tyr Asn Trp Ser Ala Lys Ala Lys
     Arg Arg Asn Thr Thr Gly Thr Gly Arg Met Arg His Leu Lys Ile
     Val Tyr Arg Arg Phe Arg His Gly Phe Arg Glu Gly Thr Thr Pro
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140
   Lys Pro Lys Arg Ala Ala Val Ala Ala Ser Ser Ser
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  Arg Thr Trp Trp Pro Arg Cys Cys Thr Thr Cys Thr His Gln Arg
  Ser Arg Trp Met Arg Arg Ala Cys Arg Ile Cys Ser Pro Arg His
  Thr Ala Ser Arg Ser Leu Pro Ser Ser Pro Ser Ala Cys Pro Ser
  Cys Arg Ser Ala Cys Ala Ser Pro Thr Ala Trp Pro Ser Ser Val
  Ser Ala Ser Cys Ser Thr Ala Arg Val Ser Pro Trp Leu Pro Ala
  Thr Ser Ser Ala Leu Thr Ser Arg Trp Trp Arg Ala Thr Leu Thr
  Ser Ser Asp Ser Arg Arg Arg Ala His Arg His His Leu Gln Arg
  Arg Ala Leu Thr Trp Arg Arg Ser Leu Val Phe Glu Ala Val
                                      115
 Met Arg Trp Ala Gly Ser Gly Asp Ala Glu Ala Glu Arg
 Gln Arg Ala Leu Pro Thr Val Phe Glu Ser Val Arg Cys Arg Leu
 Leu Pro Arg Ala Phe Leu Glu Ser Arg Val Glu Arg His Pro Leu
 Val Arg Ala Gln Pro Glu Leu Leu Arg Lys Val Gln Met Val Lys
 Asp Ala His Glu Gly Arg Ile Thr Thr Leu Arg Lys Lys Lys
 Gly Lys Asp Gly Ala Gly Ala Lys Glu Ala Asp Lys Gly Thr Ser
 Lys Ala Lys Ala Glu Glu Asp Glu Glu Ala Glu Arg Ile Leu Pro
 Gly Ile Leu Asn Asp Thr Leu Arg Phe Gly Met Phe Leu Gln Asp
Leu Ile Phe Met Ile Ser Glu Glu Gly Ala Val Ala Tyr Asp Pro
Ala Ala Asn Glu Cys Tyr Cys Ala Ser Leu Ser Ser Gln Val Pro
Lys Asn His Val Ser Leu Val Thr Lys Glu Asn Gln Val Phe Val
                                    280
Ala Gly Gly Leu Phe Tyr Asn Glu Asp Asn Lys Glu Asp Pro Met
Ser Ala Tyr Phe Leu Gln Phe Asp His Leu Val Gly Gly Gln Arg
Asp Gln Gly Arg Arg Ala Leu Pro Gly Leu Gly His Val Leu Arg
Gln Ala Val Ile Gln Met Gly
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                350
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Lys Ala Ala Val Ser Ser Phe Ser Lys Pro Leu Lys Gly Ser Ala
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                 20
Gly Gly Arg Arg Asn Ser Lys Gly Gly Pro Arg Gln Gly Ala Ile
                                      40
                 35
Gly Leu Gly Leu Arg Glu Pro Glu Thr Ala Ala Ala Ala Ala Ala
                 50
                                      55
Ala Ala Ala Gly Gly Ala Gln Gly Thr Pro Xaa Leu Pro Val Leu
                                      70
                 65
Cys Leu Gly Pro Ser Leu Leu Pro Arg Ala Gln Cys Gly Leu Ala
                 80
                                      85
Ser Val Lys Glu Phe Asp Lys Lys Tyr Asn Pro Thr Trp His Cys
                 95
                                     100
Ile Val Gly Arg Asn Phe Gly Ser Tyr Val Thr His Glu Thr Lys
                                                          120
                                     115
                110
His Phe Ile Tyr Phe Tyr Leu Gly Gln Val Ala Ile Leu Leu Phe
                125
                                     130
Lys Ser Gly
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Gly Arg Leu Ser Ser Gln Pro Trp Met Phe Ser Ser Cys Gly Arg
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Pro Ala Ser Lys Ser Thr Thr Ser Ser Thr Pro Arg Thr Arg Gln
                 20
                                      25
Leu His Ala Trp Ser Arg Cys Trp Asn Gly Ala Phe Thr Pro Cys
                                                           45
Arg Leu Ser Ala Ser Pro Ala Thr Asn Ala Thr Arg Trp Gly Met
                  50
                                      55
Ala Ala Pro Arg Cys Trp Ser Arg Pro Cys Arg Glu Thr Leu Ser
                  65
                                      70
Trp Ser Trp Arg Ala Ala Pro Trp Pro Leu Ser Pro Thr Gly Thr
                  80
                                      85
Ala Ser Trp Lys Pro Val Cys Leu Phe Pro Arg Pro Pro Gly Lys
                                     100
                                                          105
                  95
Thr Ala Pro Ala Arg Ala Val Pro Ser Arg Met Cys Ser Arg Pro
                 110
                                     115
Thr Met Gln Pro Ser Lys Ser Met Ala Pro Pro Pro Arg Arg Ala
```

				125					130					135
Leu	Pro	Leu	Pro		Val	Ala	Ser	Ala		Pro	Val	Arg	Ser	
Ser	Pro	Ala	Arg		Gln	Ala	Trp	Leu		Ala	Thr	Arg	His	
Ala	Ser	Pro	Arg		Leu	Gln	Ser	Gly		Ala	Arg	Ser	Gly	
Thr	Thr	Pro	Cys		Ala	Leu	Thr	Pro		Thr	Ala	Phe	Pro	
Val	Ala	Leu	Pro	His 200	Leu	Phe	His	Ala	Ser 205	Tyr	Trp	Glu	Ser	Thr 210
				215					220				Asp	225
				230					235				Phe	240
				245					250				His	255
_		_		260					265				Ala	270
				275					280				Met Asp	285
_			-	290					295		-		Leu	300
				305					310				Ile	315
				320			-		325		_		Met	330
			_	335	_		_		340				Val	345
	_		_	350				-	355				Ser	360
	_	_		365					370				Arg	375
Gly	Ala	Val	Asp	380 Val	Val	Arg	His	Trp	385 Gln	Asp	Leu	Gly	Tyr	390 Leu
Ile	Ile	Tyr	Val		Gly	Arg	Pro	Asp		Gln	Lys	Gln	Arg	
Val	Ala	Trp	Leu		Gln	His	Asn	Phe		His	Gly	Val	۷al	
Phe	Cys	Asp	Gly	425 Leu 440	Val	His	Asp	Pro	430 Leu 445	Arg	His	Lys	Ala	435 Asn 450
Phe	Leu	Lys	Leu		Ile	Ser	Glu	Leu		Leu	Arg	Val	His	
Ala	Tyr	Gly	Ser		Lys	Asp	Val	Ala		Tyr	Ser	Ala	Ile	
Leu	Ser	Pro	Met		Ile	Tyr	Ile	Val	Gly 490	Arg	Pro	Thr	Lys	Lys 495
Leu	Gln	Gln	Gln	Cys 500		Phe	Ile	Thr	Asp 505	Gly	Tyr	Ala	Ala	His 510
				515					520				Arg	525
Thr	Ala	Thr	Arg	Met 530		Leu	Arg	Lys	Gly 535	Ser	Phe	Gly	Leu	Pro 540
_		_	_	545		_		_	550				Arg	555
				560					565				Arg	570
	_	_		575					580		_		Met	585
val	ATG	wrg	СΤĀ	590	_	стλ	Arg	ATG	мет 595	rnr	чТĀ	Arg	Leu	600

Pro Gly Ala Ala Ala Gly Pro Lys 605

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Phe Leu Leu Ile Ile Leu Gln Arg Glu Ile Asn His Asn Arg Ala
                 380
                                     385
Leu Leu Arg Asn Asp Leu Cys Ala Leu Glu Cys Gly Ile Pro Lys
                 395
                                     400
                                                          405
His Phe Gly Leu Phe Tyr Ala Met Gly Thr Ala Leu Met Met Glu
                 410
                                     415
                                                          420
Gly Leu Leu Ser Ala Cys Tyr His Val Cys Pro Asn Tyr Thr Asn
                 425
                                     430
Phe Gln Phe Asp Thr Ser Phe Met Tyr Met Ile Ala Gly Leu Cys
                                     445
                                                          450
Met Leu Lys Leu Tyr Gln Lys Arg His Pro Asp Ile Asn Ala Ser
                 455
                                     460
Ala Tyr Ser Ala Tyr Ala Cys Leu Ala Ile Val Ile Phe Phe Ser
                 470
                                     475
Val Leu Gly Val Val Phe Gly Lys Gly Asn Thr Ala Phe Trp Ile
                 485
                                     490
Val Phe Ser Ile Ile His Ile Ile Ala Thr Leu Leu Ser Thr
                 500
                                     505
Gln Leu Tyr Tyr Met Gly Arg Trp Lys Leu Asp Ser Gly Ile Phe
                 515
                                     520
Arg Arg Ile Leu His Val Leu Tyr Thr Asp Cys Ile Arg Gln Cys
                530
                                     535
                                                          540
Ser Gly Pro Leu Tyr Val Asp Arg Met Val Leu Leu Val Met Gly
                 545
                                     550
                                                          555
Asn Val Ile Asn Trp Ser Leu Ala Ala Tyr Gly Leu Ile Met Arg
                 560
                                     565
Pro Asn Asp Phe Ala Ser Tyr Leu Leu Ala Ile Gly Ile Cys Asn
                 575
                                     580
Leu Leu Tyr Phe Ala Phe Tyr Ile Ile Met Lys Leu Arg Ser
                 590
                                     595
                                                          600
Gly Glu Arg Ile Lys Leu Ile Pro Leu Leu Cys Ile Val Cys Thr
                 605
                                     610
                                                          615
Ser Val Val Trp Gly Phe Ala Leu Phe Phe Phe Gln Gly Leu
                 620
                                     625
Ser Thr Trp Gln Lys Thr Pro Ala Glu Ser Arg Glu His Asn Arg
                 635
                                     640
Asp Cys Ile Leu Leu Asp Phe Phe Asp Asp His Asp Ile Trp His
                 650
                                     655
Phe Leu Ser Ser Ile Ala Met Phe Gly Ser Phe Leu Val Ser Gly
                 665
                                     670
                                                         675
Pro Pro Gly Arg Ala Gly Trp Val Arg Glu Gly Ser Ser Cys Leu
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Leu Leu Cys Gly
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Phe Ile Trp His Lys Ser Ile Leu Ser Arg Met Ala Glu Ala Val
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Leu Ile Asp Leu Phe Gly Leu Lys Leu Asn Ser Gln Lys Asn Cys
                 20
                                      25
His Gln Thr Leu Leu Lys Thr Leu Asn Ala Val Gln Tyr His His
                 35
                                      40
Ala Ala Lys Ala Lys Phe Leu Cys Ile Met Cys Cys Ser Asn Ile
                                      55
                                                          60
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Ser Tyr Glu Arg Asp Gly Glu Gln Asp Asn Cys Glu Ile Glu Thr
                 65
Ser Asn Gly Leu Ser Ala Leu Leu Glu Glu Phe Glu Ile Val Ser
                                                          90
                 80
                                      85
Cys Pro Ser Met Ala Ala Thr Leu Tyr Thr Ile Lys Gln Lys Ile
                                     100
                                                         105
                 95
Asp Glu Lys Asn Leu Ser Ser Ile Lys Val Ile Val Pro Arg His
                110
                                     115
Arg Lys Thr Leu Met Lys Ala Phe Ile Asp Gln Leu Phe Thr Asp
                                     130
                                                         135
                125
Val Tyr Asn Phe Glu Phe Glu Asp Leu Gln Val Thr Phe Arg Gly
                                                          150
                                     145
                140
Gly Leu Phe Lys Gln Ser Ile Glu Ile Asn Val Ile Thr Ala Gln
                                     160
                155
Glu Leu Arg Gly Ile Gln Asn Glu Ile Glu Thr Phe Leu Arg Ser
                170
                                     175
                                                          180
Leu Pro Ala Leu Arg Gly Lys Leu Thr Ile Ile Thr Ser Ser Leu
                                     190
                                                          195
                185
Ile Pro Asp Ile Phe Ile His Gly Phe Thr Thr Arg Thr Gly Gly
                                     205
                                                          210
                200
Ile Ser Tyr Ile Pro Thr Leu Ser Ser Phe Asn Leu Phe Ser Ser
                215
                                     220
                                                          225
Ser Lys Arg Arg Asp Pro Lys Val Val Gln Gly Ile Lys Thr
                                     235
                230
His His Ser Asn Asp Ile Trp Ile Met Gly Arg Lys Glu Pro Asp
                                     250
                 245
Ser Tyr Asp Gly Ile Thr Thr Asn Gln Arg Gly Val Thr Ile Ala
                260
                                     265
Ala Leu Gly Ala Asp Cys Ile Pro Ile Val Phe Ala Asp Pro Val
                                     280
                 275
Lys Lys Ala Cys Gly Val Ala His Ala Gly Trp Lys Gly Thr Leu
                                                          300
                 290
                                     295
Leu Gly Val Ala Met Ala Thr Val Asn Ala Met Ile Ala Glu Tyr
                 305
                                     310
Gly Cys Ser Leu Glu Asp Ile Val Val Val Leu Gly Pro Ser Val
                                     325
                                                          330
                 320
Gly Pro Cys Cys Phe Thr Leu Pro Arg Glu Ser Ala Glu Ala Phe
                 335
                                     340
His Asn Leu His Pro Ala Cys Val Gln Leu Phe Asp Ser Pro Asn
                 350
                                      355
Pro Cys Ile Asp Ile Arg Lys Ala Thr Ser Phe Pro Lys Asp Ser
                                                          375
                 365
                                      370
Ser Arg Thr Gly Arg Asn Ser Ser Thr Glu Tyr Ser Gly Pro Glu
                                                          390
                 380
                                     385
Pro Arg Ser Gln Pro Leu Tyr Ile Leu Pro Ser
                 395
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Ser Gln His Phe Pro Arg Pro Ser Val Glu Thr Glu Val Gly Asp
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                                       10
Tyr Met Phe Cys Phe Asp Asn Thr Phe Ser Thr Ile Ser Glu Lys
                                       25
                                                           30
                  20
Val Ile Phe Phe Glu Leu Ile Leu Asp Asn Met Gly Glu Gln Ala
                                                           45
                  35
                                       40
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Gln Glu Gln Glu Asp Trp Lys Lys Tyr Ile Thr Gly Thr Asp Ile
                 50
                                      55
Leu Asp Met Lys Leu Glu Asp Ile Leu Glu Ser Ile Asn Ser Ile
                 65
                                      70
                                                           75
Lys Ser Arg Leu Ser Lys Ser Gly His Ile Gln Thr Leu Leu Arg
                 80
                                      85
Ala Phe Glu Ala Arg Asp Arg Asn Ile Gln Glu Ser Asn Phe Asp
                 95
                                     100
Arg Val Asn Phe Trp Ser Met Val Asn Leu Val Val Met Val Val
                110
                                     115
                                                        . 120
Val Ser Ala Ile Gln Val Tyr Met Leu Lys Ser Leu Phe Glu Asp
                125
Lys Arg Lys Ser Arg Thr
                140
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Ala Trp Ala Ala Val Ile Thr Pro Trp Asn Phe Pro Ser Ala Met
Ile Thr Arg Lys Val Gly Ala Ala Leu Ala Ala Gly Cys Thr Val
                 20
                                      25
Val Val Lys Pro Ala Glu Asp Thr Pro Phe Ser Ala Leu Ala Leu
                                      40
                                                           45
Ala Glu Leu Ala Ser Gln Ala Gly Ile Pro Ser Gly Val Tyr Asn
                 50
                                      55
Val Ile Pro Cys Ser Arg Lys Asn Ala Lys Glu Val Gly Glu Ala
                 65
                                      70
Ile Cys Thr Asp Pro Leu Val Ser Lys Ile Ser Phe Thr Gly Ser
                 80
                                      85
Thr Thr Thr Gly Lys Ile Leu Leu His His Ala Ala Asn Ser Val
                 95
                                     100
Lys Arg Val Ser Met Glu Leu Gly Gly Leu Ala Pro Phe Ile Val
                110
                                     115
Phe Asp Ser Ala Asn Val Asp Gln Ala Val Ala Gly Ala Met Ala
                125
                                     130
Ser Lys Phe Arg Asn Thr Gly Gln Thr Cys Val Cys Ser Asn Gln
                140
                                     145
                                                          150
Phe Leu Val Gln Arg Gly Ile His Asp Ala Phe Val Lys Ala Phe
                155
                                     160
Ala Glu Ala Met Lys Lys Asn Leu Arg Val Gly Asn Gly Phe Glu
                170
                                     175
                                                          180
Glu Gly Thr Thr Gln Gly Pro Leu Ile Asn Glu Lys Ala Val Glu
                185
                                                          195
Lys Val Glu Lys Gln Val Asn Asp Ala Val Ser Lys Gly Ala Thr
                200
                                     205
Val Val Thr Gly Gly Lys Arg His Gln Leu Gly Lys Asn Phe Phe
                215
                                     220
Glu Pro Thr Leu Leu Cys Asn Val Thr Gln Asp Met Leu Cys Thr
                230
                                     235
His Glu Glu Thr Phe Gly Pro Leu Ala Pro Val Ile Lys Phe Asp
                245
                                     250
                                                          255
Thr Glu Glu Glu Ala Ile Ala Ile Ala Asn Ala Ala Asp Val Gly
                260
                                     265
                                                          270
Leu Ala Gly Tyr Phe Tyr Ser Gln Asp Pro Ala Gln Ile Trp Arg
                275
                                     280
                                                          285
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Val Ala Glu Gln Leu Glu Val Gly Met Val Gly Val Asn Glu Gly
                                     295
Leu Ile Ser Ser Val Glu Cys Pro Phe Gly Gly Val Lys Gln Ser
                290
                                     310
Gly Leu Gly Arg Glu Gly Ser Lys Tyr Gly Ile Asp Glu Tyr Leu
                320
Glu Leu Lys Tyr Val Cys Tyr Gly Gly Leu
                335
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 Lys Leu Asp Tyr Arg Arg Glu Pro Pro Arg Pro Val Tyr Val Leu
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 Trp Ile Cys Glu Tyr Ser Tyr Ser Val Leu Phe Ile Asn Thr Tyr
                                       25
 Asp Leu Thr Gln Lys Val Lys Val Asn Thr Leu Trp Gly Gly Pro
 Val Ser Val Gln Gly Gly Ser Pro Ala Arg Lys Gly Cys Ser Leu
                   35
 Arg Cys His Ser Ser Phe Ser Pro Ala Ser Asp His Ile Cys His
  Ser Gly Pro Glu Gly Ala Gly Gly Pro Ser Asn Gln Ala Arg Ser
                   65
  Trp Ser Arg Gln Gly Gly Phe Arg Gly Phe Gly Ala Ala Phe Val
  Ser Arg Cys Arg Gln Lys Leu Gln Phe Ser Ser Val Cys Phe Val
                   95
                                       115
  Ser Ser Ala Arg Arg Ser Pro Ala Cys Val Ala Leu Arg Pro Ala
                  110
                                       130
  Gly Ile Gly Arg Ser Thr Ala Lys Thr Pro Gly Pro Pro Gly Ser
                                       145
  Leu Glu Met Gly Ala Leu Thr Phe Arg Asp Val Ala Ile Glu Phe
   Ser Leu Glu Glu Trp Gln Cys Leu Asp Thr Glu Gln Gln Asn Leu
                   155
                   170
   Tyr Arg Asn Val Met Leu Asp Asn Tyr Arg Asn Leu Val Phe Leu
   Gly Ile Ala Val Ser Lys Pro Asp Leu Ile Thr Cys Leu Glu Gln
   Glu Lys Glu Pro Trp Asn Leu Lys Thr His Asp Met Val Ala Lys
                   215
   Pro Pro Gly Arg
   <210> 136
   <211> 407
    <212> PRT
    <213> Homo sapiens
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    <221> misc_feature
    <223> Incyte ID No: LG:144920.1.orf1:2002JAN18
    Val Leu Leu Asp Glu Ala Gln Arg Leu Leu Tyr Arg Asp Val Met
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10
  Leu Glu Asn Phe Ala Leu Met Ala Ser Leu Gly Cys Trp His Gly
                                        25
  Met Glu Asp Glu Glu Ile Pro Phe Glu Gln Ser Phe Ser Ile Gly
                   35
                                        40
  Met Ser Gln Ile Arg Ile Pro Lys Gly Gly Pro Ser Thr Gln Lys
                                        55
  Ala Tyr Pro Cys Gly Thr Cys Gly Leu Val Leu Lys Asp Ile Leu
                                        70
  His Leu Ala Glu His Gln Glu Thr His Pro Gly Gln Lys Pro Tyr
                   80
                                        85
 Met Cys Val Leu Cys Gly Lys Gln Phe Trp Phe Ser Ala Asn Leu
                   95
                                       100
 His Gln His Gln Lys Gln His Ser Gly Glu Lys Pro Phe Arg Ser
                  110
                                      115
 Asp Lys Ser Arg Pro Phe Leu Leu Asn Asn Cys Ala Val Gln Ser
                  125
                                      130
 Leu Glu Met Ser Phe Val Thr Gly Glu Ala Cys Lys Asp Phe Leu
                                      145
 Ala Ser Ser Ser Ile Phe Glu His His Ala Pro His Asn Glu Trp
                  155
                                      160
 Lys Pro His Ser Asn Thr Lys Cys Glu Glu Ala Ser His Cys Gly
                  170
 Lys Arg His Tyr Lys Cys Ser Glu Cys Gly Lys Thr Phe Ser Arg
                                      175
                  185
                                      190
 Lys Asp Ser Leu Val Gln His Gln Arg Val His Thr Gly Glu Arg
                 200
                                      205
 Pro Tyr Glu Cys Gly Glu Cys Gly Lys Thr Phe Ser Arg Lys Pro
                 215
                                      220
 Ile Leu Ala Gln His Gln Arg Ile His Thr Gly Glu Met Pro Tyr
                 230
                                      235
 Glu Cys Gly Ile Cys Gly Lys Val Phe Asn His Ser Ser Asn Leu
                 245
                                      250
 Ile Val His Gln Arg Val His Thr Gly Ala Arg Pro Tyr Lys Cys
                 260
                                     265
Ser Glu Cys Gly Lys Ala Tyr Ser His Lys Ser Thr Leu Val Gln
                 275
                                     280
His Glu Ser Ile His Thr Gly Glu Arg Pro Tyr Glu Cys Ser Glu
                 290
                                     295
Cys Gly Lys Tyr Phe Gly His Lys Tyr Arg Leu Ile Lys His Trp
                                                          300
                 305
                                     310
Ser Val His Thr Gly Ala Arg Pro Tyr Glu Cys Ile Ala Cys Gly
                 320
                                     325
Lys Phe Phe Ser Gln Ser Ser Asp Leu Ile Ala His Gln Arg Val
                335
                                     340
His Asn Gly Glu Lys Pro Tyr Val Cys Ser Glu Cys Gly Lys Ala
                350
                                     355
Phe Ser His Lys His Val Leu Val Gln His His Arg Ile His Thr
                365
                                     370
Gly Glu Arg Pro Tyr Lys Cys Ser Glu Cys Gly Lys Ala Phe Arg
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                                     385
Gln Arg Ala Ser Leu Ile Arg His Trp Lys Ile His Thr Gly Glu
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Arg Pro
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<211> 529

<212> PRT

<213> Homo sapiens

<220>

<221> misc_feature

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Leu Pro Gln Glu Phe Gly Asp Gln Asp Leu Leu Gln Met Phe Met
                  455
                                      460
 Pro Phe Gly Asn Val Val Ser Ala Lys Val Phe Ile Asp Lys Gln
                 470
                                      475
                                                           480
 Thr Asn Leu Ser Lys Cys Phe Gly Phe Val Ser Tyr Asp Asn Pro
                 485
                                      490
 Val Ser Ala Gln Ala Ala Ile Gln Ser Met Asn Gly Phe Gln Ile
                 500
                                      505
                                                           510
 Gly Met Lys Arg Leu Lys Val Gln Leu Lys Arg Ser Lys Asn Asp
                                      520
 Ser Lys Pro Tyr
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Gly His Gly Thr Asp Cys Gly Glu Cys His Arg Arg His Thr His
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                                      25
                                                           30
Arg Pro Val Phe Gln Ser Ser His Tyr Thr Val Asn Val Asn Glu
                                      40
Asp Arg Pro Ala Gly Thr Thr Val Val Leu Ile Ser Ala Thr Asp
                  50
                                                           60
Glu Asp Thr Gly Glu Asn Ala Arg Ile Thr Tyr Phe Met Glu Asp
                  65
Ser Ile Pro Gln Phe Arg Ile Asp Ala Asp Thr Gly Ala Val Thr
                  80
                                      85
Thr Gln Ala Glu Leu Asp Tyr Glu Asp Gln Val Ser Tyr Thr Leu
                  95
                                     100
Ala Ile Thr Ala Arg Asp Asn Gly Ile Pro Gln Lys Ser Asp Thr
                 110
                                     115
Thr Tyr Leu Glu Ile Leu Val Asn Asp Val Asn Asp Asn Ala Pro
                 125
                                     130
                                                          135
Gln Phe Leu Arg Asp Ser Tyr Gln Gly Ser Val Tyr Glu Asp Val
                 140
                                     145
                                                          150
Pro Pro Phe Thr Ser Val Leu Gln Ile Ser Ala Thr Asp Arg Asp
                 155
                                     160
                                                          165
Ser Gly Leu Asn Gly Arg Val Phe Tyr Thr Phe Gln Gly Gly Asp
                 170
                                     175
Asp Gly Asp Gly Asp Phe Ile Val Glu Ser Thr Ser Gly Ile Val
                185
                                     190
                                                          195
Arg Thr Leu Arg Arg Leu Asp Arg Glu Asn Val Ala Gln Tyr Val
                 200
                                     205
                                                          210
Leu Arg Ala Tyr Ala Val Asp Lys Gly Met Pro Pro Ala Arg Thr
                215
                                     220
                                                          225
Pro Met Glu Val Thr Val Thr Val Leu Asp Val Asn Asp Asn Pro
                230
                                     235
Pro Val Phe Glu Gln Asp Glu Phe Asp Val Phe Val Glu Glu Asn
                245
                                     250
Ser Pro Ile Gly Leu Ala Val Ala Arg Val Thr Ala Thr Asp Pro
                260
                                     265
                                                          270
Asp Glu Gly Thr Asn Ala Gln Ile Met Tyr Gln Ile Val Glu Gly
                275
                                     280
Asn Ile Pro Glu Val Phe Gln Leu Asp Ile Phe Ser Gly Glu Leu
                290
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300

Thr	Ala	Leu	Val	Asp 305	Leu	Asp	Tyr	Glu	Asp 310	Arg	Pro	Glu	Tyr	Val 315
Leu	Val	Ile	Gln	Ala 320	Thr	Ser	Ala	Pro	Leu 325	Val	Ser	Arg	Ala	Thr 330
Val	His	Val	Arg	Leu 335	Leu	Asp	Arg	Asn	Asp 340.		Pro	Pro	Val	Leu 345
Gly	Asn	Phe	Glu	Ile 350	Leu	Phe	Asn	Asn	Tyr 355	Val	Thr	Asn	Arg	Ser 360
				365					370				His	375
				380					385				Gly	390
				395					400				Leu	405
				410					415				Ile	420
				425					430				Gln	435
				440					445				His	450
			-	455					460				Leu -	465
				470					475				Leu	480
				485					490				Asp	495
_				500					505				Val	510
				515	٠.				520				Ser	525
				530					535				Thr	540
				545					550				Cys	555
				560					565				Leu	570
	_			575					580				Leu Pro	585
				590					595				Cys	600
				605					610				Glu	615
				620					625					630 Cys
				635					640				Суз	645
				650					655				Lys	660
				665					670				. Val	675
				680	I				685				Gly	690
				695	ı				700				Thr	705
				710	1				715					720 Lys
				725	,				730	1				735 Leu
				740	)				745	,				750 Val
				755	;				760	)				765 Lys

				770					775					780
Tyr	Tyr	Asn	Lys	Pro 785	Leu	Leu	Gly	Gln	Thr 790	Gly	Leu	Pro	Gln	Gly 795
Pro	Ser	Glu	Gln	Lys 800	Val	Ala	Val	Val	Thr 805	Val	Asp	Gly	Cys	Asp 810
Thr	Gly	Val	Ala	Leu 815	Arg	Phe	Gly	Ser	Val 820	Leu	Gly	Asn	Tyr	Ser 825
Cys	Ala	Ala	Gln	Gly 830	Thr	Gln	Gly	Gly	Ser 835	Lys	Lys	Ser	Leu	Asp '840
Leu	Thr	Gly	Pro	Leu 845	Leu	Leu	Gly	Gly	Val 850	Pro	Asp	Leu	Pro	Glu 855
Ser	Phe	Pro	Val	Arg 860	Met	Arg	Gln	Phe	Val 865	Gly	Cys	Met	Arg	Asn 870
				Ser 875				_	880					885
				Val 890			•		895					900
				Cys 905					910	_				915
_				Cys 920		_			925		_	_	_	930
				Met 935					940					945
			_	His 950	_				955					960
				Met 965					970		_	_		975
				Thr 980			-		985					990
				Val 995				:	1000				:	1005
			;	Arg 1010 Gln				:	1015					1020
			:	1025 Ser				:	1030				:	1035
				1040 Arg					1045				:	1050
			;	1055 Pro				:	1060				:	1065
			:	1070 Gln					1075					1080
			:	1085 Asp				:	1090				:	1095
			:	1100 Leu					1105				:	1110
_				1115 Cys					1120				;	1125
			:	1130 Tyr				:	1135				;	1140
				1145 Glu	_		_	:	1150					1155
			;	1160 Tyr				• :	1165				:	1170
			:	1175 Thr				:	1180				:	1185
			;	1190 Thr			_	:	1195				:	1200
				1205 Asp					1210					1215
				1220 Tyr				:	1225					1230
				1235	- 3				1240			-,,,		1245

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Cys Asp Cys Tyr Pro Thr Gly Ser Leu Ser Arg Val Cys Asp Pro Glu Asp Gly Gln Cys Pro Cys Lys Pro Gly Val Ile Gly Arg Gln Cys Asp Arg Cys Asp Asn Pro Phe Ala Glu Val Thr Thr Asn Gly Cys Glu Val Asn Tyr Asp Ser Cys Pro Arg Ala Ile Glu Ala Gly Ile Trp Trp Pro Arg Thr Arg Phe Gly Leu Pro Ala Ala Ala Pro Cys Pro Lys Gly Ser Phe Gly Thr Ala Val Arg His Cys Asp Glu His Arg Gly Trp Leu Pro Pro Asn Leu Phe Asn Cys Thr Ser Ile Thr Phe Ser Glu Leu Lys Gly Phe Ala Glu Arg Leu Gln Arg Asn Ser Gly Arg Ser Gln Gln Leu Ala Leu Leu 1355 Glu Ser Gly Leu Asp 1375 Leu Arg Asn Ala Thr Gln His Thr Ala Gly Tyr Phe Gly Ser Asp Val Lys Val Ala Tyr Gln Leu Ala Thr Arg Leu Leu Ala His Glu 1405 Ser Thr Gln Arg Gly Phe Gly Leu Ser Ala Thr Gln Asp Val His Phe Thr Glu Asn Leu Leu Arg Val Gly Ser Ala Leu Leu Asp Thr Ala Asn Lys Arg His Trp Glu Leu Ile Gln Gln Thr Glu Gly Gly Thr Ala Trp Leu Leu Gln His Tyr Glu Ala Tyr Ala Ser Ala Leu 1465 Ala Gln Asn Met Arg His Thr Tyr Leu Ser Pro Phe Thr Ile Val Thr Pro Asn Ile Val Ile Ser Val Val Arg Leu Asp Lys Gly Asn 1495 Phe Ala Gly Ala Lys Leu Pro Arg Tyr Glu Ala Leu Arg Gly Glu 1490 Gln Pro Pro Asp Leu Glu Thr Thr Val Ile Leu Pro Glu Ser Val Phe Arg Glu Thr Pro Pro Val Val Arg Pro Ala Gly Pro Gly Glu Ala Gln Glu Pro Glu Glu Leu Ala Arg Arg Gln Arg Arg His Pro Glu Leu Ser Gln Gly Glu Ala Val Ala Ser Val Ile Ile Tyr Arg Thr Leu Ala Gly Leu Leu Pro His Asn Tyr Asp Pro Asp Lys Arg Ser Leu Arg Val Pro Lys Arg Pro Ile Ile Asn Thr Pro Val Val Ser Ile Ser Val His Asp Asp Glu Glu Leu Leu Pro Arg Ala Leu Asp Lys Pro Val Thr Val Gln Phe Arg Leu Leu Glu Thr Glu Glu Arg Thr Lys Pro Ile Cys Val Phe Trp Asn His Ser Ile Leu Val Ser Gly Thr Gly Gly Trp Ser Ala Arg Gly Cys Glu Val Val Phe Arg Asn Glu Ser His Val Ser Cys Gln Cys Asn His Met Thr Ser Phe Ala Val Leu Met Asp Val Ser Arg Arg Glu Asn Gly Glu Ile Leu Pro Leu Lys Thr Leu Thr Tyr Val Ala Leu Gly Val Thr Leu Ala Ala Leu Leu Thr Phe Phe Phe Leu Thr Leu Leu Arg Ile

1715
Leu Arg Ser Asn Gln His Gly Ile Arg Arg Asn Leu Thr Ala Ala
Leu Gly Leu Ala Gln Leu Val Phe Leu Leu Gly Ile Asn Gln Ala
Asp Leu Pro Phe Ala Cys Thr Val Ile Ala Ile Leu Leu His Phe
1760 1765 Leu Tyr Leu Cys Thr Pho Court 1765 1770
1775 The Ser Trp Ala Leu Leu Glu Ala Leu His
Led Tyr Arg Ala Leu Thr Glu Val Arg Asp Val Asn Thr Gly Pro
Met Arg Phe Tyr Tyr Met Leu Gly Trp Gly Val Pro Ala Phe Ile
Thr Gly Leu Ala Val Gly Leu Asp Pro Glu Gly Tyr Gly Asp Pro
Asp Phe Cys Trp Leu Ser Ile Tyr Asp Thr Leu Il
Ala Gly Pro Val Ala Phe Ala Val Ser Wet Ser Val Phe Leu Tyr
1850 1855 1860
Ile Leu Ala Ala Arg Ala Ser Cys Ala Ala Gln Arg Gln Gly Phe 1865 1870 1875
Glu Lys Lys Gly Pro Val Ser Gly Leu Gln Pro Ser Phe Ala Val
1895 Leu Leu Ser Ala Thr Trp Leu Leu Ala Leu Leu Ser Val
1910 Ash Sel Asp Thr Leu Leu Phe His Tyr Leu Phe Ala Thr Cys Asn
Cys Ile Gln Gly Pro Phe Ile Phe Leu Ser Tyr Val Val Leu Ser
Lys Glu Val Arg Lys Ala Leu Lys Leu Ala Cys Sor Arg 1935
Ser Pro Asp Pro Ala Leu Thr Thr Lys Ser Thr Leu Thr Ser Ser
1955 1960 1965  Tyr Asn Cys Pro Ser Pro Tyr Ala Asp Gly Arg Leu Tyr Gln Pro 1970
1970 1975 Leu Tyr Gln Pro Tyr Gly Asp Ser Mag Clu G
Tyr Gly Asp Ser Ala Gly Ser Leu His Ser Thr Ser Arg Ser Gly  1985 1990 1995
Lys Ser Gln Pro Ser Tyr Ile Pro Phe Leu Leu Arg Glu Glu Ser  2000 2005 2010
Ald Leu Asn Pro Gly Gln Gly Pro Pro Gly Leu Gly Asp Pro Gly
Ser Leu Phe Leu Glu Gly Gln Asp Gln Gln His Asp Pro Asp Thr
Asp Ser Asp Ser Asp Leu Ser Leu Glu Asp Asp Gln Ser Clu Gan
Tyr Ala Ser Thr His Ser Ser Asp Ser Glu
Glu Glu Glu Glu Ala Ala Phe Pro Gly Glu Gln Gly Trp Asp Ser
2075 2080 2085
Leu Leu Gly Pro Gly Ala Glu Arg Leu Pro Leu His Ser Thr Pro 2090 2095 2100
Lys Asp Gly Gly Pro Gly Pro Gly Lys Ala Pro Trp Pro Gly Asp 2110  Pho Cly The Tile 2115
2120 2120 2120
Glu Arg Leu Arg Glu Asn Gly Asp Ala Leu Ser Arg Glu Gly Ser
Leu Gly Pro Leu Pro Gly Ser Ser Ala Gln Pro His Lys Cly Ile
Leu Lys Lys Cys Leu Pro Thr Ile Ser Glu Lys Ser Cor Leu
Leu Arg Leu Pro Leu Glu Gln Cys Thr Gly Ser Ser Arg Gly Ser
2180 2185 Ser Ser Arg Gly Ser 2190

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Ser Ala Ser Glu Gly Ser Arg Gly Gly Pro Pro Pro Arg Pro Pro
               2195
                                    2200
Pro Arg Gln Ser Leu Gln Glu Gln Leu Asn Gly Val Met Pro Ile
                                    2215
                                                         2220
               2210
Ala Met Ser Ile Lys Ala Gly Thr Val Asp Glu Asp Ser Ser Gly
                                    2230
               2225
Ser Glu Phe Leu Phe Phe Asn Phe Leu His
                                    2245
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Glu Pro Arg Tyr Pro Gly Phe Val Leu Gly Leu Asp Val Gly Ser
                                      25
                 20
Ser Val Ile Arg Cys His Val Tyr Asp Arg Ala Ala Arg Val Cys
                                                           45
                 35
                                      40
Gly Ser Ser Val Gln Lys Val Glu Asn Leu Tyr Pro Gln Ile Gly
                                      55
                 50
Trp Val Glu Ile Asp Pro Asp Val Leu Trp Ile Gln Phe Val Ala
                                                           75
                 65
Val Ile Lys Glu Ala Val Lys Ala Ala Gly Ile Gln Met Asm Gln
                  80
                                      85
Ile Val Gly Leu Gly Ile Ser Thr Gln Arg Ala Thr Phe Ile Thr
                 95
                                     100
Trp Asn Lys Lys Thr Gly Asn His Phe His Asn Phe Ile Ser Trp
                110
                                     115
                                                          120
Gln Asp Leu Arg Ala Val Glu Leu Val Lys Ser Trp Asn Asn Ser
                                    . 130
                                                          135
                125
Leu Leu Met Lys Ile Phe His Ser Ser Cys Arg Val Leu His Phe
                140
                                     145
Phe Thr Arg Ser Lys Arg Leu Phe Thr Ala Ser Leu Phe Thr Phe
                 155
                                      160
Thr Thr Gln Gln Thr Ser Leu Arg Leu Val Trp Ile Leu Gln Asn
                 170
                                      175
Leu Thr Glu Val Gln Lys Ala Val Glu Glu Glu Asn Cys Cys Phe
                                      190
                 185
Gly Thr Ile Asp Thr Trp Leu Leu Tyr Lys Leu Thr Lys Gly Ser
                                                          210
                 200
                                      205
Val Tyr Ala Thr Asp Phe Ser Asn Ala Ser Thr Thr Gly Leu Phe
                 215
Asp Pro Tyr Lys Met Cys Trp Ser Gly Met Ile Thr Ser Leu Ile
                 230
                                      235
Ser Ile Pro Leu Ser Leu Leu Pro Pro Val Arg Asp Thr Ser His
                                      250
                 245
Asn Phe Gly Ser Val Asp Glu Glu Ile Phe Gly Val Pro Ile Pro
                                      265
                                                          270
                 260
Ile Val Ala Leu Val Ala Asp Gln Gln Ser Ala Met Phe Gly Glu
                                      280
                                                          285
                 275
Cys Cys Phe Gln Thr Gly Asp Val Lys Leu Thr Met Gly Thr Gly
                                                           300
                 290
                                      295
Thr Phe Leu Asp Ile Asn Thr Gly Asn Ser Leu Gln Gln Thr Thr
                                                          315
                 305
                                      310
Gly Gly Phe Tyr Pro Leu Ile Gly Trp Lys Ile Gly Gln Glu Val
                                      325
                 320
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Val Cys Leu Ala Glu Ser Asn Ala Gly Asp Thr Gly Thr Ala Ile
                335
                                     340
Lys Trp Ala Gln Gln Leu Asp Leu Phe Thr Asp Ala Ala Glu Thr
                350
                                     355
Glu Lys Met Ala Lys Ser Leu Glu Asp Ser Glu Gly Val Cys Phe
                365
                                     370
                                                         375
Val Pro Ser Phe Ser Gly Leu Gln Ala Pro Leu Asn Asp Pro Trp
                                     385
                                                         390
Ala Cys Ala Ser Phe Met Gly Leu Lys Pro Ser Thr Ser Lys Tyr
                395
                                     400
His Leu Val Arg Ala Ile Leu Glu Ser Ile Ala Phe Arg Asn Lys
                410
                                     415
                                                         420
Gln Leu Tyr Glu Met Met Lys Lys Glu Ile His Ile Pro Val Arg
                425
                                     430
Lys Ile Arg Ala Asp Gly Gly Val Cys Lys Asn Gly Phe Val Met
                440
                                     445
Gln Met Thr Ser Asp Leu Ile Asn Glu Asn Ile Asp Arg Pro Ala
                455
                                     460
                                                         465
Asp Ile Asp Met Ser Cys Leu Gly Ala Ala Ser Leu Ala Gly Leu
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                                     475
Ala Val Gly Phe Gly Leu Thr Arg Arg Asn
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Ala Leu Ser Arg Gly Ala Glu Leu Arg Val Pro Gly Gly Ala Lys
His Gly Met Cys Leu Leu Gly Ala Thr Gly Val Gly Lys Thr
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Leu Leu Val Lys Arg Leu Gln Glu Val Ser Ser Arg Asp Gly Lys
                 35
                                      40
Gly Asp Leu Gly Glu Pro Pro Pro Thr Arg Pro Thr Val Gly Thr
                 50
                                      55
Asn Leu Thr Asp Ile Val Ala Gln Arg Lys Ile Thr Ile Arg Glu
                 65
                                      70
Leu Gly Gly Cys Met Gly Pro Ile Trp Ser Ser Tyr Tyr Gly Asn
                 80
                                      85
Cys Arg Ser Leu Leu Phe Val Met Asp Ala Ser Asp Pro Thr Gln
                 95
                                     100
Leu Ser Ala Ser Cys Val Gln Leu Leu Gly Leu Leu Ser Ala Glu
                110
                                     115
                                                         120
Gln Leu Ala Glu Ala Ser Val Leu Ile Leu Phe Asn Lys Ile Asp
                                     130
Leu Pro Cys Tyr Met Ser Thr Glu Glu Met Lys Ser Leu Ile Arg
                140
                                     145
                                                         150
Leu Pro Asp Ile Ile Ala Cys Ala Lys Gln Asn Ile Thr Thr Ala
                155
                                     160
                                                         165
Glu Ile Ser Ala Arg Glu Gly Thr Gly Leu Ala Gly Val Leu Ala
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                                     175
Trp Leu Gln Ala Thr His Arg Ala Asn Asp
                185
                                     190
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Leu Ser Thr Tyr Glu Leu Thr Leu Glu Arg Lys Pro Tyr Glu Cys
Lys Val Cys Gly Lys Ala Phe Thr Thr Ser Ser His Leu Ile Val
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                                      25
His Ile Arg Ser His Thr Gly Glu Lys Pro Tyr Ile Cys Lys Glu
                                      40
Cys Gly Lys Ala Phe Ala Ser Ser Ser His Leu Ile Glu His Arg
                                      55
                 50
Arg Thr His Thr Gly Glu Lys Pro Tyr Ile Cys Asn Glu Cys Gly
                 65
                                      70
Lys Ala Phe Arg Ala Ser Ser His Leu His Lys His Gly Arg Ile
                                      85
                 80
His Thr Gly Gln Lys Pro Tyr Lys Cys Lys Glu Cys Gly Lys Ala
                                     100
                                                          105
                 95
Tyr Asn Arg Phe Tyr Leu Leu Lys Glu His Leu Lys Thr Tyr Thr
                                                          120
                110
                                     115
Glu Glu Gln Val Phe Val Cys Lys Asp Cys Gly Lys Ser Phe Lys
                                     130
                                                          135
                125
Asn Ser Ser Cys Leu Asn His His Thr Gln Ile His Thr Asp Glu
                                                          150
                                     145
                140
Lys Pro Phe
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<222> (1) ... (608)
<223> unknown or other
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Gly Pro Gly Ser Asp Pro Arg Ala Ser Ser Gln Thr Met Arg Arg
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Arg Ser Trp Asp Gln Ala Cys Leu Leu Gln Glu Lys Gln Glu Glu
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                                      25
Gly Lys Asp Pro Glu Gly Gln Pro Leu Leu Ala Pro Gln Arg Val
                                       40
                  35
Arg Ser Gly Ala Ala Ala Xaa Leu Gln Gln Val Arg Thr Lys Glu
                                       55
 Cys Trp Ser Trp Glu Ser Tyr Leu Glu Glu Gln Lys Ala Ile Thr
                                       70
                  65
 Ala Pro Val Ser Leu Phe Gln Asp Ser Gln Ala Val Thr His Asn
                  80
                                       85
 Lys Asn Gly Phe Lys Leu Gly Met Lys Leu Glu Gly Ile Asp Pro
                                                           105
                  95
                                      100
 Gln His Pro Ser Met Tyr Phe Ile Leu Thr Val Ala Glu Val Cys
                                                           120
                                      115
                 1.10
 Gly Tyr Arg Leu Arg Leu His Phe Asp Gly Tyr Ser Glu Cys His
                 125
                                      130
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As	p Ph	e Tr	p Va	1 Ası 14(	n Ala	a Ası	n Se	r Pr	o Asp 145	] Ile	His	s Pro	Ala	Gly
Tr	p Ph	e Gl	u Ly		Gly	/ His	5 Ly:	s Le	u Glr 160	ı Pro	Pro	) Lys	Gl ₂	150 Tyr
Ly	s Gl	u Gl	u Gl	u Phe 170	e Sei	Tr	Se	c Gli	n Tyr 175	Leu	Arg	g Ser	Thi	165 Arg
Ala	a Gl	n Al	a Ala	a Pro	Lys	His	Let	ı Phe	e Val 190	Ser	Glr	ı Ser	His	180 Ser
•				200	,				y Met	Lys				195 Val 210
				2T	)				val 220	Ala				Asp
				230	ļ				Phe 235	Asp				Asp
				240	)				250	ı				Pro
				Gln 260	,				265					Gln
				275	1				280					Glu
				Ser 290					295					200
				305					310					215
				Ala 320					325					220
				335					340					215
				Ile 350					355					260
				Lys 365 Ser					370					275
				380 Leu					485					200
				395 Cys					400					40E
				410 Phe					415					400
				425 Asn					430					40 C
				440 Ala					445					150
				455 Arg					ΛΕΛ					
				Pro					475					400
				Ser					49ก					405
				Ser					505 Gln					E10
				Gly					520					FOF
				Phe					535 Thr					E 40
				Leu					550 Met					
				Thr					565 Val					~~~
				575 Ala 590					580 Asn					
				Asp					595					600

605

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Leu Ser His Val Pro Ser Ala Ala Ala Gln Gly Ala Trp Ser Trp
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                                      25
Glu Trp Tyr Leu Lys Glu Gln Lys Ala Val Ala Pro Val Glu
                 35
                                      40
Leu Phe Ser Lys Asp Gln Ser Phe Pro Glu His Glu Asn Gly Phe
                 50
                                      55
Gln Ile Gly Met Arg Leu Glu Gly Ile Asp Pro Arg His Pro Ser
                 65
                                      70
Val Phe Cys Val Leu Ser Val Ala Glu Val Cys Gly Tyr Arg Leu
                 80
                                      85
Arg Leu His Phe Asp Gly Tyr Leu Ser Cys Tyr Asp Phe Trp Thr
                 95
                                     100
Asn Ala Gly Ser Pro Asp Ile His Pro Val Gly Trp Cys Glu Lys
                110
                                     115
                                                         120
Thr Lys His Glu Leu His Ile Pro Lys Gly Tyr Arg Lys Asp Lys
                 125
                                     130
                                                         135
Phe Val Trp Met Asp Tyr Leu Lys Ala Cys Lys Leu Gln Asn Ala
                140
                                     145
Pro Lys Lys Leu Phe Arg Asn Arg Ser Pro Asn Gly Pro Met Ser
                155
                                     160
Lys Glu Phe Gln Val Gly Met Lys Leu Glu Ala Val Asp Arg Lys
                170
                                     175
Asn Pro Ser Leu Val Cys Val Ala Thr Ile Ala Asp Ile Val Glu
                185
                                     190
                                                          195
Asp Arg Leu Leu Val His Phe Asp Asn Trp Gly Asp Ser Tyr Asp
                 200
                                     205
Tyr Trp Cys Asp Val Asn Ser Pro Tyr Val Gln Pro Val Gly Trp
                215
                                     220
                                                          225
Cys Gln Glu Asn Gly Arg Thr Leu Ile Ala Pro Gln Gly Tyr Pro
                 230
                                     235
Ile Gln Lys Ile Phe Pro Gly Gln Asn Thr Trp Lys Leu Leu Lys
                 245
                                     250
                                                          255
Pro Met Gln Phe Leu Pro Lys Phe Leu Lys
                 260
                                     265
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Tyr Gln Arg Gln Leu Lys Glu Met Asn Phe Glu Thr Ser Arg Cys
Ala Thr Leu Gln Tyr Cys Pro Asp Pro Tyr Ile Gln Arg Phe Val
                  20
                                      25
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Glu	Thr	Pro	Ala	His	Phe	Ser	Trp	Lys	Glu 40		Туг	Туг	Arg	Ser 45
Thr	Met	Ser	Gln	Ser 50	Thr	Gln	Thr	Asn		Phe	Lev	Ser	Pro	Glu 60
Val	Phe	Gln	His	Ile 65	Trp	Asp	Phe	Leu		Gln	Pro	Il∈	Cys	Ser 75
Val	Gln	Pro	Ile	Asp 80	Leu	Asn	Phe	Val	Asp 85		Pro	Ser	Glu	Asp 90
Gly	Ala	Thr	Asn	Lys 95	Ile	Glu	Ile	Ser		Asp	Суз	Ile	Arg	Met 105
Gln	Asp	Ser	Asp	Leu 110	Ser	Asp	Pro	Met	Trp		Gln	Туг	Thr	Asn 120
				125					130				Gly	Ser 135
				140					145				Ser	Val 150
				155					160				Asp	Ala
				170					175				Pro	Gly 180
				185					190				Ala	195
				200					205				Tyr	210
				215					220				Thr	225
				230					235				Lys	240
				245					250				His	255
				260					265				Ser	270
				275					280				Glu	285
				290					295				Pro	300
•				305					310				Met	315
				320					325				Leu	330
				335					340				Arg	345
				350					355				Arg Ser	360
				365					370				Ser	375
				380					385				Asp	390
				395					400				Glu	405
				410					415				Leu	420
				425					430				Gln	435
				440					445				Ser	450
				455					460					465
				470					475				Arg	480
				485					490				Asn	495
							_	- 1			1			

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510
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Pro Met Met Gly Thr His Met Pro Met Ala Gly Asp Met Asn Gly
                                    520
                515
Leu Ser Pro Thr Gln Ala Leu Pro Pro Pro Leu Ser Met Pro Ser
                530
                                     535
Thr Ser His Cys Thr Pro Pro Pro Pro Tyr Pro Thr Asp Cys Ser
                                                         555
                                     550
                545
Ile Val Ser Phe Leu Ala Arg Leu Gly Cys Ser Ser Cys Leu Asp
                                     565
                560
Tyr Phe Thr Thr Gln Gly Leu Thr Thr Ile Tyr Gln Ile Glu His
                                     580
                575
Tyr Ser Met Asp Asp Leu Ala Ser Leu Lys Ile Pro Glu Gln Phe
                                                          600
                                     595
                590
Arg His Ala Ile Trp Lys Gly Ile Leu Asp His Arg Gln Leu His
                                     610
                605
Glu Phe Ser Ser Pro Ser His Leu Leu Arg Thr Pro Ser Ser Ala
                                                          630
                                     625
                620
Ser Thr Val Ser Val Gly Ser Ser Glu Thr Arg Gly Glu Arg Val
                                     640
                635
Ile Asp Ala Val Arg Phe Thr Leu Arg Gln Thr Ile Ser Phe Pro
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Pro Arg Asp Glu Trp Asn Asp Phe Asn Phe Asp Met Asp Ala Arg
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Arg Asn Lys Gln Gln Arg Ile Lys Glu Glu Gly Glu
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 Val Tyr Pro Asp Glu Leu Pro Asn Thr Ser Val Val Ile Val Phe
                                       25
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 His Asn Glu Ala Trp Ser Thr Leu Leu Arg Thr Val Tyr Ser Val
                                       40
                  35
 Ile Asn Arg Ser Pro His Tyr Leu Leu Ser Glu Val Ile Leu Val
                                       55
                  50
 Asp Asp Ala Ser Glu Arg Asp Phe Leu Lys Leu Thr Leu Glu Asn
                  65
 Tyr Val Lys Asn Leu Glu Val Pro Val Lys Ile Ile Arg Met Glu
                                       85
                  80
 Glu Arg Ser Gly Leu Ile Arg Ala Arg Leu Arg Gly Ala Ala Ala
                                      100
                  95
 Ser Lys Gly Gln Val Ile Thr Phe Leu Asp Ala His Cys Glu Cys
                                      115
                 110
 Thr Leu Gly Trp Leu Glu Pro Leu Leu Ala Arg Ile Lys Glu Asp
                                                           135
                                      130
                 125
 Arg Lys Thr Val Val Cys Pro Ile Ile Asp Val Ile Ser Asp Asp
                                      145
                 140
 Thr Phe Glu Tyr Met Ala Gly Ser Asp Met Thr Tyr Gly Gly Phe
                                                           165
                                      160
                  155
 Asn Trp Lys Leu Asn Phe Arg Trp Tyr Pro Val Pro Gln Arg Glu
                                                           180
                                      175
                  170
 Met Asp Arg Arg Lys Gly Asp Arg Thr Leu Pro Val Arg Thr Pro
                                      190
                  185
 Thr Met Ala Gly Gly Leu Phe Ser Ile Asp Arg Asn Tyr Phe Glu
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200
                                      205
 Glu Ile Gly Thr Tyr Asp Ala Gly Met Asp Ile Trp Gly Gly Glu
                 215
                                      220
                                                          225
 Asn Leu Glu Met Ser Phe Arg Ile Trp Gln Cys Gly Gly Ser Leu
                 230
                                      235
 Glu Ile Val Thr Cys Ser His Val Gly His Val Phe Arg Lys Ala
                 245
                                      250
 Thr Pro Tyr Thr Phe Pro Gly Gly Thr Gly His Val Ile Asn Lys
                 260
                                      265
                                                          270
Asn Asn Arg Arg Leu Ala Glu Val Trp Met Asp Glu Phe Lys Asp
                 275
                                      280
                                                          285
 Phe Phe Tyr Ile Ile Ser Pro Gly Val Val Lys Val Asp Tyr Gly
                 290
                                      295
                                                          300
Asp Val Ser Val Arg Lys Thr Leu Arg Glu Asn Leu Lys Cys Lys
                 305
                                      310
                                                          315
Pro Phe Ser Trp Tyr Leu Glu Asn Ile Tyr Pro Asp Ser Gln Ile
                 320
                                      325
Pro Arg Arg Tyr Tyr Ser Leu Gly Glu Ile Arg Asn Val Glu Thr
                 335
                                      340
Asn Gln Cys Leu Asp Asn Met Gly Arg Lys Glu Asn Glu Lys Val
                 350
                                      355
Gly Ile Phe Asn Cys His Gly Met Gly Gly Asn Gln Val Phe Ser
                 365
                                      370
                                                          375
Tyr Thr Ala Asp Lys Glu Ile Arg Thr Asp Asp Leu Cys Leu Asp
                 380
                                      385
                                                          390
Val Ser Arg Leu Asn Gly Pro Val Ile Met Leu Lys Cys His His
                 395
                                      400
Met Arg Gly Asn Gln Leu Trp Glu Tyr Asp Ala Glu Thr His Thr
                 410
                                      415
                                                          420
Leu Leu His Ile Ile Thr Gln Ser Cys Leu Ser Val Asn Lys Val
                 425
                                      430
                                                          435
Ala Asp Gly Ser Gln His Pro Thr Val Glu Thr Cys Asn Asp Ser
                 440
                                     445
Thr Leu Gln Lys Trp Leu Leu Arg Asn Tyr Thr Arg Met Glu Ile
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Phe Arg Asn Ile Phe Gly Asn Ser Thr Asp Tyr Ile Leu
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Val Met Phe Ala Asp Arg Lys Ser Lys Leu Ser Arg Glu Phe Arg
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Ala Leu Gly Thr Cys Tyr Leu Asp Gln Phe Glu Ala Ser Leu Ile
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Tyr Trp Thr Asn Trp Gln Gln Lys Met Lys Ser Ser Val Ala Gln
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Ile Lys Pro Ser Ser Gly His Asp Arg Arg Glu Asn Leu Asn Pro
                 50
                                      55
Tyr Gln Arg Asn Ser Ser Pro Glu Asp Arg Tyr Glu Glu Glu Glu
                 65
                                      70
Arg Ser Pro Arg Asp Arg Asp Tyr Phe Asp Tyr Ser Arg Ser Asp
                 80
                                      85
Tyr Glu His Ser Arg Arg Gly Arg Ser Tyr Asp Ser Ser Met Glu
                 95
                                     100
Ser Arg Asn Arg Asp Arg Glu Lys Arg Arg Glu Arg Glu Arg Asp
```

		Arg	<b>T</b>						115					
				125					Pro 130					135
Asn	Pro	Glu	Thr	Ser 140	Val	Thr	Gln	Ser	Ser 145	Ser	Ala	Gln	Asp	Glu 150
Pro	Ala	Thr	Lys	Lys 155	Lys	Lys	Asp	Glu	Leu 160	Asp	Pro	Leu	Leu	Thr 165
Arg	Thr	Gly	Gly	Ala 170	Tyr	Ile	Pro	Pro	Ala 175	Lys	Leu	Arg	Met	Met 180
				185					Leu 190					195
				200					Asn 205					210
				215					Ile 220					225
				230					Leu 235					240
				245					Thr 250					255
				260					Pro 265					270
				275					Arg 280					285
				290					Ser 295					300
				305					Val 310					315
				320					Asp 325					330
				335					Leu 340					345
				350					Glu 355					360
				365					Val ·370					375
				380					Phe 385					390
				395					Glu 400					405
				410					Asn 415					420
				425					Met 430					435
_	_			440					445					Asp 450
				455					460					Glu 465
				470	1				475					Val 480
				485	,				490					495
				500	)				505					Cys 510
				515	,				520					Lys 525
				530	)				535					Thr 540
				545	,				550	ļ.				555
				560	)				565	;				570
Ту	c Asp	Thr	: Ile	His 575		, Leu	ı Glu	Thr	Asn 580	Lys	Leu	ı Arç	, Asn	585

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Ala Lys Met Phe Ala His Leu Leu Tyr Thr Asp Ser Leu Pro Trp
                 590
                                    595
 Ser Val Leu Glu Cys Ile Lys Leu Ser Glu Glu Thr Thr Ser
                 605
                                    610
                                                        615
 Ser Ser Arg Ile Phe Val Lys Ile Phe Phe Gln Glu Leu Cys Glu
                620
                                    625
 Tyr Met Gly Leu Pro Lys Leu Asn Ala Arg Leu Lys Asp Glu Thr
                635
                                    640
                                                        645
 Leu Gln Pro Phe Phe Glu Gly Leu Leu Pro Arg Asp Asn Pro Arg
                650
                                    655
Asn Thr Arg Phe Ala Ile Asn Phe Phe Thr Ser Ile Gly Leu Gly
                665
                                    670
Gly Leu Thr Asp Glu Leu Arg Glu His Leu Lys Asn Thr Pro Lys
                680
                                    685
                                                        690
Val Ile Val Ala Gln Lys Pro Asp Val Glu Gln Asn Lys Ser Ser
                695
                                    700
710
                                    715
Ser Asp Ser Asp Ser Asp Ser Ser Ser Ser Glu Ser Ser
                725
                                    730
Ser Glu Glu Ser Asp Ser Ser Ser Ile Ser Ser His Ser Ser Ala
                740
                                    745
Ser Ala Asn Asp Val Arg Lys Lys Gly His Gly Lys Thr Arg Ser
                755
                                    760
Lys Glu Val Asp Lys Leu Ile Arg Asn Gln Gln Thr Asn Asp Arg
                770
                                    775
                                                        780
Lys Gln Lys Glu Arg Arg Gln Glu His Gly His Gln Glu Thr Arg
                785
                                    790
Thr Glu Arg Glu Arg Arg Ser Glu Lys His Arg Asp Gln Asn Ser
                800
                                    805
Arg Gly Ser Asn Trp Arg Asp Pro Ile Thr Lys Tyr Thr Ser Asp
                815
                                    820
Lys Asp Val Pro Ser Glu Arg Asn Asn Tyr Ser Arg Val Ala Asn
                830
                                    835
Asp Arg Asp Gln Glu Met His Ile Asp Leu Glu Asn Lys His Gly
                845
                                    850
Asp Pro Lys Lys Arg Gly Glu Arg Arg Asn Ser Phe Ser Glu
                860
                                    865
Asn Glu Lys His Thr His Arg Ile Lys Asp Ser Glu Asn Phe Arg
                875
                                    880
Arg Lys Asp Arg Ser Lys Ser Lys Glu Met Asn Arg Lys His Ser
                890
                                   895
Gly Ser Arg Ser Asp Glu Asp Arg Tyr Gln Asn Gly Ala Glu Arg
                905
                                   910
Arg Trp Glu Lys Ser Ser Arg Tyr Ser Glu Gln Ser Arg Glu Ser
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                                   925
                                                       930
Lys Lys Asn Gln Asp Arg Arg Glu Lys Ser Pro Ala Lys Gln
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                                   940
Lys
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PCT/US03/01363 WO 03/062379

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Gly Ala Ala Leu Arg Asp Val Leu Gly Arg Ala Gln Gly Val Leu
                                      40
                 35
Phe Asp Cys Asp Gly Val Leu Trp Asn Gly Glu Arg Ala Val Pro
                 50
Gly Ala Pro Glu Leu Leu Glu Arg Leu Ala Arg Ala Gly Lys Ala
Ala Leu Phe Val Ser Asn Asn Ser Arg Arg Ala Arg Pro Glu Leu
                 80
                                      85
Ala Leu Arg Phe Ala Arg Leu Gly Phe Gly Gly Leu Arg Ala Glu
                 95
                                     100
Gln Leu Phe Ser Ser Ala Leu Cys Ala Ala Arg Leu Leu Arg Gln
                                                         120
                                    115
                110
Arg Leu Pro Gly Pro Pro Asp Ala Pro Gly Ala Val Phe Val Leu
                                                         135
                125
                                    130
Gly Gly Glu Gly Leu Arg Ala Glu Leu Arg Ala Ala Gly Leu Arg
                                     145
                140
Leu Ala Gly Asp Pro Ser Ala Gly Asp Gly Ala Ala Pro Arg Val
                155
                                     160
Arg Ala Val Leu Val Gly Tyr Asp Glu His Phe Ser Phe Ala Lys
                170
                                    175
Leu Arg Glu Ala Cys Ala His Leu Arg Asp Pro Glu Cys Leu Leu
                185
                                     190
Val Ala Thr Asp Arg Asp Pro Trp His Pro Leu Ser Asp Gly Ser
                200
                                     205
Arg Thr Pro Gly Thr Gly Ser Leu Ala Ala Ala Val Glu Thr Ala
                215
                                     220
Ser Gly Arg Gln Ala Leu Val Val Gly Lys Pro Ser Pro Tyr Met
                                     235
                230
Phe Glu Cys Ile Thr Glu Asn Phe Ser Ile Asp Pro Ala Arg Thr
                                     250
                245
Leu Met Val Gly Asp Arg Leu Glu Thr Asp Ile Leu Phe Gly His
                                     265
                260
Arg Cys Gly Met Thr Thr Val Leu Thr Leu Thr Gly Val Ser Arg
                275
                                     280
Leu Glu Glu Ala Gln Ala Tyr Leu Ala Ala Gly Gln His Asp Leu
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                290
Val Pro His Tyr Tyr Val Glu Ser Ile Ala Asp Leu Thr Glu Gly
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                 305
Leu Glu Asp
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<213> Homo sapiens

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Val	Ser	Thr	Pro	Gly 80	Pro	Ser	Arg	Lys	Arg 85		Arg	r Leu	Ser	Asn 90
				95					100				Tyr	Asn
				110					115				Cys	Glu 120
				125					130				Tyr	Tyr 135
				140					145				Gln	. 150
				155					160				Ala	165
				170					175				Pro	180
				185					190				Tyr	195
				200					205				Asp	210
				215					220				Thr	225
				230					235				Leu	240
				245					250				Ile	255
				260					265				Asp Ala	270
				275					280				Ala	285
				290					295				Glu	300
				305					310				Ile	315
				320					325				Lys	330
				335					340				Arg	345
				350					355				Arg	360
				365 Tyr					370				Gly	375
				380 Ile					385				Glu	390
				395 Ala					400				Gly	405
				410 Ser					415 Met				Asn	420
Phe	Tyr	Gln	Leu	425 Tyr 440	Asp	Thr	Val	Arg	430 Ser	Cys	Arg	Asn	Asn	435 Gln
Gly	Gln	Leu	Ile	Ala 455	Glu	Pro	Phe	Tyr	445 His	Leu	Pro	Ser	Lys	
Lys	Tyr	Pro	Asp		Tyr	Gln	Gln	Ile	460 Lys	Met	Pro	Ile	Ser	
Gln	Gln	Ile	Arg		Lys	Leu	Lys	Asn	475 Gln 490	Glu	Tyr	Glu	Thr	
Asp	His	Leu	Glu		Asp	Leu	Asn	Leu	Met 505	Phe	Glu	Asn	Ala	
Arg	Tyr	Asn	Val		Asn	Ser	Ala	Ile	Tyr 520	Lys	Arg	Val	Leu	510 Lys 525
				Met 530					Lys 535				Arg	Arg 540
Asp	Asp	Ile	Glu	Asp	Gly	Asp	Ser	Met	Ile	Ser	Ser	Ala	Thr	Ser

				545					550					555
Asp	Thr	Gly	Ser	Ala 560	Lys	Arg	Lys	Ser	Lys 565	Lys	Asn	Ile	Arg	Lys 570
Gln	Arg	Met	Lys	Ile 575	Leu	Phe	Asn	Val	Val 580	Leu	Glu	Ala	Arg	Glu 585
Pro	Gly	Ser	Gly	Arg 590	Arg	Leu	Cys	Asp	Leu 595	Phe	Met	Val	Lys	Pro 600
Ser	Lys	Lys	Asp	Тут 605	Pro	Asp	Tyr	Tyr	Lys 610	Ile	Ile	Leu	Glu	Pro 615
Met	Asp	Leu	Lys	Ile 620	Ile	Glu	His	Asn	Ile 625	Arg	Asn	Asp	Lys	Туr 630
	-			635			Glu		640					645
				650			Glu.		655					660
				665			Leu		670					675
_				680			Asp		685					690
				695			Ser		700					705
				710			Asn		715					720
	_			725			Arg		730					735
_				740			Leu		745					11e 750
				755			Lys		760					765
				770			Ser		775					780
				785			Tyr		790		,			795
				800			His		-805					810
				815			Asp		820					825
				830			His		835					840
				845			Arg		850					855
				860			Asn		865					870
				875			Asn		880					885
				890					895					Arg 900
				905					910					Ala 915
				920					925					930
				935					940					Thr 945
				950					955					<b>Leu</b> 960
				965					970	)				975
				980	ı				985					990
				995					1000	1				1005
Ser	Met	Tyr	His	Val 1010		' Asp	Tyr	· Val	Tyr 1015	val	GLU	Pro	) Ala	Glu 1020

				1025	)			1	ነበገለ				Trp Glu 1035
				1040	}			ı Tyr 1	Gly	Cys			Tyr Arg
				T022	)			1	L060				Glu Lys
				T0.00	)			1	L075				Ser Lys
				1085	i			1	L090				Phe Lys
				1100	)		_	1	1.05				Cys Glu 1110
				TTT2	1			1	.120				Lys Leu 1125
				1130				1	135				Asp Val 1140
				1145				1	150				Ala Asp 1155
				<b>TTPO</b>				1	165				Ser Arg 1170
				TT/2				1	.180				Val Pro
				<b>TT30</b>				1	195				Glu Gln 1200 Val Phe
				1205				1	210				1215 Ile Glu
				T220				1	225				1230 Pro Ile
				1235				1	240				1245 Met Phe
				T220				1	255				1260 Cys Pro
				1265				1	270				1275 Lys Asp
				T580				1	285				1290 Ile Leu
			Ser	Arg				· 1:	300				1305 Lys Lys
			Leu	FÄ2				Leu :	315 Ser				1320 Val Asp
Asp	Glu	Ile	Tyr	1325 Tyr	Phe	Arg	Lys	Pro :	330 Ile	Val	Pro	Gln	1335 Lys Glu
Pro	Ser	Pro	Leu	1340 Leu	Glu	Lys	Lys	Ile	345 Gln	Leu	Leu	Glu.	1350 Ala Lys
Phe	Ala	Glu	Leu	1355 Glu 1370	Gly	Gly	Asp	Asp A	360 Asp	Ile	Glu	Glu	1365 Met Gly
Glu	Glu	Asp	Ser	Glu 1385	Val	Ile	Glu	Pro 1	375 Pro 390	Ser	Leu	Pro	1380 Gln Leu
Gln	Thr	Pro	Leu	Ala 1400	Ser	Glu	Leu	Asp I	Leu 405	Met	Pro	Tyr	1395 Thr Pro 1410
Pro	Gln	Ser	Thr		Lys	Ser	Ala	Lys (	3ly 420	Ser	Ala	Lys	Lys Glu 1425
			J	L430				Ser 0	31y 135				Phe Ser
			Arg	Ala 1445				Ala (	3ln 150				Tyr Ser
			J	L460				Gly 7	Chr 165				Asn Leu 1470
			1	4/5				Glu G	3ly 180				Gln Gly
Val	Ala	Pro	Met	Val	Gly	Thr	Pro	Ala	Pro	Gly	Gly	Ser	Pro Tyr

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1495
               1490
Gly Gln Gln Val Gly Val Leu Gly Pro Pro Gly Gln Gln Ala Pro
               1505
                                   1510
Pro Pro Tyr Pro Gly Pro His Pro Ala Gly Pro Pro Val Ile Gln
                                    1525
               1520
Gln Pro Thr Thr Pro Met Phe Val Ala Pro Pro Pro Lys Thr Gln
                                    1540
               1535
Arg Leu Leu His Ser Glu Ala Tyr Leu Lys Tyr Ile Glu Gly Leu
                                    1555
               1550
Ser Ala Glu Ser Asn Ser Ile Ser Lys Trp Asp Gln Thr Leu Ala
                                    1570
               1565
Ala Arg Arg Arg Asp Val His Leu Ser Lys Glu Gln Glu Ser Arg
                                    1585
                                                        1590
               1580
Leu Pro Ser His Trp Leu Lys Ser Lys Gly Ala His Thr Thr Met
                                    1600
               1595
                                                        1605
Ala Asp Ala Leu Trp Arg Leu Arg Asp Leu Met Leu Arg Asp Thr
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Leu Asn Ile Arg Gln Ala Tyr Asn Leu Glu Asn Val
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Ala Ala Glu Arg Arg Ala Pro Glu Glu Glu Leu Pro Pro Leu Asp
                  35
                                      40
Pro Glu Glu Ile Arg Lys Arg Leu Glu His Thr Glu Arg Gln Phe
                                      55
                  50
Arg Asn Arg Arg Lys Ile Leu Ile Arg Gly Leu Pro Gly Asp Val
                                      70
                  65
Thr Asn Gln Glu Val His Asp Leu Leu Ser Asp Tyr Glu Leu Lys
                                      85
                  80
Tyr Cys Phe Val Asp Lys Tyr Lys Gly Thr Ala Phe Val Thr Leu
                                     100
                  95
Leu Asn Gly Glu Gln Ala Glu Ala Ala Ile Asn Ala Phe His Gln
                                                          120
                                     115
                 110
Ser Arg Leu Arg Glu Arg Glu Leu Ser Val Gln Leu Gln Pro Thr
                                      130
                 125
Asp Ala Leu Leu Cys Val Ala Asn Leu Pro Pro Ser Leu Thr Gln
                 140
                                      145
Gln Gln Phe Glu Glu Leu Val Arg Pro Phe Gly Ser Leu Glu Arg
                                      160
                 155
Cys Phe Leu Val Tyr Ser Glu Arg Thr Gly Gln Ser Lys Gly Tyr
                                      175
                                                          180
                 170
Gly Phe Ala Glu Tyr Met Lys Lys Asp Ser Ala Ala Arg Ala Lys
                                                          195
                 185
                                      190
 Ser Asp Leu Leu Gly Lys Pro Leu Gly Pro Arg Thr Leu Tyr Val
                                      205
                                                          210
                 200
His Trp Thr Asp Ala Gly Gln Leu Thr Pro Ala Leu Leu His Ser
                                                          225
                                      220
                 215
 Arg Cys Leu Cys Val Asp Arg Leu Pro Pro Gly Phe Asn Asp Val
                 230
                                      235
 Asp Ala Leu Cys Arg Ala Leu Ser Ala Val His Ser Pro Thr Phe
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				245					250					255
Cys	Gln	Leu	Ala	Cys 260	Gly	Gln	Asp	Gly	250 Gln 265	Leu	Lys	Gly	Phe	255 Ala 270
Val	Leu	Glu	Tyr	Glu 275	Thr	Ala	Glu	Met	Ala 280	Glu	Glu	Ala	Gln	
Gln	Ala	Asp	Gly	Leu 290	Ser	Leu	Gly	Gly	-	His	Leu	Arg	Val	
Phe	Cys	Ala	Pro	Gly 305	Pro	Pro	Gly	Arg	Ser	Met	Leu	Ala	Ala	
Ile	Ala	Ala	Gln	Ala 320	Thr	Ala	Leu	Asn	Arg 325	Gly	Lys	Gly	Leu	
Pro	Glu	Pro	Asn	Ile 335	Leu	Gln	Leu	Leu	Asn 340	Asn	Leu	Gly	Pro	Ser 345
Ala	Ser	Leu	Gln	Leu 350	Leu	Leu	Asn	Pro	Leu 355	Leu	His	Gly	Ser	Ala 360
				Gly 365			_		370					375
				Ala 380					385					390
				Gln 395	-			_	400		_	_		405
				Gln 410					415					420
				Ala 425					430					435
_	_	_	<u>.</u>	Pro 440					445				-	450
				Arg 455					460					465
				Pro 470			-		475					480
				Gln 485					490					495
				Leu 500 Leu					505					510
				515 Glu					520					525
				530 Glu					535					540
				545 Ser					550					555
				560 Ser					565					570
				575 Ser					580				_	585
				590 His					595					600
				605 Lys					610					615
Arg	Ser	Ser	Gly	620 Gly	Ser	Gly	Gly	Gly	625 Pro	Leu	Ser	His	Phe	630 Tyr
Ser	Gly	Ser	Pro	635 Thr	Ser	Tyr	Phe	Thr	640 Ser	Gly	Leu	Gln	Ala	
Leu	Lys	Gln	Ser	650 His	Leu	Ser	Lys	Ala		Gly	Ser	Ser	Pro	660 Leu
Gly	Ser	Gly	Glu	665 Gly	Leu	Leu	Gly	Leu		Pro	Gly	Pro	Asn	675 Gly
His	Ser	His	Leu	680 Leu	Lys	Thr	Pro	Leu		Gly	Arg	Asn	Ala	
Leu	Pro	Thr	Cys	695 Cys	Pro	Arg	Pro	Ser		Ala	Gln	Lys	Ala	
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Met Trp Ala Ser Thr Pro Arg Pro Arg Arg His Tyr Ala Asp Ser
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Thr
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 Leu Lys Glu Val Val Phe Ser Pro Gly Leu Leu Leu Thr Ser Leu
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 Phe Ser Gln Cys Ile Trp Val Val Ser Phe Leu Ser Ser Phe Phe
                  20
                                       40
                  35
 Leu Ser Leu Pro Tyr Gly Val Ala Val Gly Val Ala Phe Ser Val
                                       55
 Leu Val Val Phe Gln Thr Gln Phe Arg Asn Gly Tyr Ala Leu
                  50
                                       70
 Ala Gln Val Met Asp Thr Asp Ile Tyr Val Asn Pro Lys Thr Tyr
                  65
                   80
 Asn Arg Ala Gln Asp Ile Gln Gly Ile Lys Ile Ile Thr Tyr Cys
                                      100
                   95
  Ser Pro Leu Tyr Phe Ala Asn Ser Glu Ile Phe Arg Gln Lys Val
                                      115
  Ile Ala Lys Thr Val Ser Leu Gln Glu Leu Gln Gln Asp Phe Glu
                  110
                                      130
  Asn Ala Pro Pro Thr Asp Pro Asn Asn Asn Gln Thr Pro Ala Asn
                  125
                                      -145
  Gly Thr Ser Val Ser Tyr Ile Thr Phe Ser Pro Asp Ser Ser Ser
                  140
                                       160
                  155
  Pro Ala Gln Ser Glu Pro Pro Ala Ser Ala Glu Ala Pro Gly Glu
                                       175
  Pro Ser Asp Met Leu Ala Ser Val Pro Pro Phe Val Thr Phe His
                  170
                                       190
  Thr Leu Ile Leu Asp Met Ser Gly Val Ser Phe Val Asp Leu Met
                  185
                                       205
                   200
  Gly Ile Lys Ala Leu Ala Lys Leu Ser Ser Thr Tyr Gly Lys Ile
                                       220
                   215
   Gly Val Lys Val Phe Leu Val Asn Ile His Ala Gln Val Tyr Asn
                                       235
                   230
   Asp Ile Ser His Gly Gly Val Phe Glu Asp Gly Ser Leu Glu Cys
                                       250
                   245
   Lys His Val Phe Pro Ser Ile His Asp Ala Val Leu Phe Ala Gln
                                        265
                   260
   Ala Asn Ala Arg Asp Val Thr Pro Gly His Asn Phe Gln Gly Ala
                                        280
                   275
   Pro Gly Asp Ala Glu Leu Ser Leu Tyr Asp Ser Glu Glu Asp Ile
                                        295
                   290
   Arg Ser Tyr Trp Asp Leu Glu Glu Met Phe Gly Ser Met Phe
                                        310
                   305
   His Ala Glu Thr Leu Thr Ala Leu
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Lys Pro Val Leu Ala Asp Gly Arg Lys Leu Pro Pro Tyr Ile Ile
                425
                                     430
Leu Arg Gly Thr Tyr Ile Pro Pro Gly Lys Phe Pro Ser Gly Met
                                                         450
                                     445
                440
Glu Ile Arg Cys His Arg Tyr Gly Trp Met Thr Glu Asp Leu Met
                                     460
                455
Gln Asp Trp Leu Glu Val Val Trp Arg Arg Arg Thr Gly Ala Val
                                     475
                470
Pro Lys Gln Arg Gly Met Leu Ile Leu Asn Gly Phe Arg Gly His
                                     490
                485
Gly Lys Asp Ser Val Lys Asn Ser Met Glu Ser Met Asn Thr Asp
                                     505
                500
Met Val Ile Ile Pro Gly Gly Leu Thr Ser Gln Leu Gln Val Leu
                                     520
                                                         525
                515
Asp Val Val Tyr Lys Pro Leu Asn Asp Ser Val Arg Ala Gln
                                     535
                                                         540
                530
Tyr Ser Asn Trp Leu Leu Ala Gly Asn Leu Ala Leu Ser Pro Thr
                545
                                     550
                                                         555
Gly Asn Ala Lys Lys Pro Pro Leu Gly Leu Phe Leu Glu Trp Val
                                                         570
                                     565
                560
Met Val Ala Trp Asn Ser Ile Ser Ser Glu Ser Ile Val Gln Gly
                                     580
                575
Phe Lys Lys Cys His Ile Ser Ser Asn Leu Glu Glu Glu Asp Asp
                590
                                     595
                                                          600
Val Leu Trp Glu Ile Glu Ser Glu Leu Pro Gly Gly Glu Pro
                                     610
                ·605
Pro Lys Asp Cys Asp Thr Glu Ser Met Ala Glu Ser Asn
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Asn Lys Gly Lys Thr Ser Leu Ala His Gln Phe Val Glu Gly Glu
                                      25
                 20
Phe Ser Glu Gly Tyr Asp Pro Thr Val Glu Asn Thr Tyr Ser Lys
                                                           45
                  35
                                      40
Ile Val Thr Leu Gly Lys Asp Glu Phe His Leu His Leu Val Asp
                                                           60
                 50
                                      55
Thr Ala Gly Gln Asp Glu Tyr Ser Ile Leu Pro Tyr Ser Phe Ile
                  65
Ile Gly Val His Gly Tyr Val Leu Val Tyr Ser Val Thr Ser Leu
                  80
                                      85
His Ser Phe Gln Val Ile Glu Ser Leu Tyr Gln Lys Leu His Glu
                                     100
                 95
Gly His Gly Lys Thr Arg Val Pro Val Val Leu Val Gly Asn Lys
                                     115
                                                          120
                 110
Ala Asp Leu Ser Pro Glu Arg Glu Val Gln Ala Val Glu Gly Lys
                                                          135
                                     130
                 125
Lys Leu Ala Glu Ser Trp Gly Ala Thr Phe Met Glu Ser Ser Ala
                 140
                                      145
Arg Glu Asn Gln Leu Thr Gln Gly Ile Phe Thr Lys Val Ile Gln
                 155
                                      160
                                                          165
Glu Ile Ala Arg Val Glu Asn Ser Tyr Gly Gln Glu Arg Arg Cys
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His Leu Met

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Arg Arg Pro Pro Ala Arg Ala Arg Gln Gly Lys Trp Asp Ala Arg
                                      25
                  20
Lys Gly Pro Leu Ala Pro Ala His His Ser Gly Ser His Ser Glu
                                      40
                  35
Tyr Pro Met Ser Ser Ser Gly Leu Pro Cys Ser Trp Trp Trp Thr
                                      55
                  50
Gln Ala Thr Pro Ala Pro Thr Trp Thr Thr Ser Ser Arg Leu Ala
                  65
                                      70
Arg Ala Pro Arg Ala Ser Cys Ala Ser Pro Pro Cys Ala Ala Arg
                                                           90
                  80
                                      85
Ala Ser Trp Trp Pro Ser Arg Arg Trp Thr Cys Ala Ser Ser Arg
                                                          105
                                     100
                  95
Gly Arg Glu Leu Leu Phe Asn Glu Val Val Ile Met Arg Asp Tyr
                                                          120
                                     115
                 110
 Gln His Glu Asn Val Val Glu Met Tyr Asn Ser Tyr Leu Val Gly
                                                          135
                                     130
                 125
 Asp Glu Leu Trp Val Val Met Glu Phe Leu Glu Gly Gly Ala Leu
                 140
 Thr Asp Ile Val Thr His Thr Arg Met Asn Glu Glu Gln Ile Ala
                 155
                                     160
 Ala Val Cys Leu Ala Val Leu Gln Ala Leu Ser Val Leu His Ala
                                      175
                 170
 Gln Gly Val Ile His Arg Asp Ile Lys Ser Asp Ser Ile Leu Leu
                                                          195
                 185
                                      190
 Thr His Asp Gly Arg Val Lys Leu Ser Asp Phe Gly Phe Cys Ala
                                      205
                 200
 Gln Val Ser Lys Glu Val Pro Arg Arg Lys Ser Leu Val Gly Thr
                                                           225
                                      220
                 215
 Pro Tyr Trp Met Ala Pro Glu Leu Ile Ser Arg Leu Pro Tyr Gly
                                      235
                  230
 Pro Glu Val Asp Ile Trp Ser Leu Gly Ile Met Val Ile Glu Met
                                      250
                 245
 Val Asp Gly Glu Pro Pro Tyr Phe Asn Glu Pro Pro Leu Lys Ala
                 260
                                      265
 Met Lys Met Ile Arg Asp Asn Leu Pro Pro Arg Leu Lys Asn Leu
                 275
                                      280
 His Lys Val Ser Pro Ser Leu Lys Gly Phe Leu Asp Arg Leu Leu
                                      295
                  290
 Val Arg Asp Pro Ala Gln Arg Ala Thr Ala Ala Glu Leu Leu Lys
                                      310
 His Pro Phe Leu Ala Lys Ala Gly Pro Pro Ala Ser Ile Val Pro
                  320
                                      325
 Leu Met Arg Gln Asn Arg Thr Arg
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 <212> PRT
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Gly Ala Met Ala Gly Val Gly Ala Gly Pro Leu Arg Ala Met Gly
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Arg Gln Ala Leu Leu Leu Leu Ala Leu Cys Ala Thr Gly Ala Gln
Gly Leu Tyr Phe His Ile Gly Glu Thr Glu Lys Arg Cys Phe Ile
                                      40
Glu Glu Ile Pro Asp Glu Thr Met Val Ile Gly Asn Tyr Arg Thr
                 35
                                      55
Gln Met Trp Asp Lys Gln Lys Glu Val Phe Leu Pro Ser Thr Pro
                 50
                  65
Gly Leu Gly Met His Val Glu Val Lys Asp Pro Asp Gly Lys Met
                                      85
 Leu Gln Val Val Leu Ser Arg Gln Tyr Gly Ser Glu Gly Arg Phe
                                     100
 Thr Phe Thr Ser His Thr Pro Gly Asp His Gln Ile Cys Leu His
                  95
                                     115
 Ser Asn Ser Thr Arg Met Ala Leu Phe Ala Gly Gly Lys Leu Arg
                 110
                                     130
 Val His Leu Asp Ile Gln Val Gly Glu His Ala Asn Asn Tyr Pro
                 125
 Glu Ile Ala Ala Lys Asp Lys Leu Thr Glu Leu Gln Leu Arg Ala
                 140
                                      160
                 155
 Arg Gln Leu Leu Asp Gln Val Glu Gln Ile Gln Lys Glu Gln Asp
                                      175
                 170
 Tyr Gln Arg Ala Ser Ala Tyr Leu Leu Val Ile
                                      190
                  185
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  Ala Ala Val Ala Phe Gly Ala Glu Val Gly Val Arg Leu Ala
  Leu Phe Ala Ala Phe Leu Val Thr Glu Leu Leu Pro Pro Phe Gln
                                        25
  Arg Leu Ile Gln Pro Glu Glu Met Trp Leu Tyr Arg Asn Pro Tyr
                   20
                                        40
  Val Glu Ala Glu Tyr Phe Pro Thr Lys Pro Met Phe Lys Ala Asp
                                        55
                    50
   Thr Arg Asp Ser Arg Gln Ala Cys Leu Ala Ala Ser Leu Ala Leu
                                        70
                    65
   Ala Leu Asn Gly Val Phe Thr Asn Thr Ile Lys Leu Ile Val Gly
                                        85
                    80
   Arg Pro Arg Pro Asp Phe Phe Tyr Arg Cys Phe Pro Asp Gly Leu
                                       100
                    95
   Ala His Ser Asp Leu Met Cys Thr Gly Asp Lys Asp Val Val Asn
                                        115
                   110
   Glu Gly Arg Lys Ser Phe Pro Ser Gly His Ser Ser Phe Ala Phe
                                        130
                   125
   Ala Gly Leu Ala Phe Ala Ser Phe Tyr Leu Ala Gly Lys Leu His
```

```
140
                                       145
  Cys Phe Thr Pro Gln Gly Arg Gly Lys Ser Trp Arg Phe Cys Ala
                  155
                                       160
  Phe Leu Ser Pro Leu Leu Phe Ala Ala Val Ile Ala Leu Ser Arg
                  170
                                       175
  Thr Cys Asp Tyr Lys His His Trp Gln Asp Val Leu Val Gly Ser
                  185
                                       190
  Met Ile Gly Met Thr Phe Ala Tyr Val Cys Tyr Arg Gln Tyr Tyr
                  200
                                       205
  Pro Pro Leu Thr Asp Ala Glu Cys His Lys Pro Phe Gln Asp Lys
                                                           210
                  215
                                       220
  Leu Val Leu Ser Thr Ala Gln Lys Pro Gly Asp Ser Tyr Cys Phe
                  230
                                       235
 Asp Asn Leu Lys Ile Glu Ser Gly Arg Ala Trp Trp Leu Met Pro
                  245
                                       250
 Val Ile Pro Thr Leu Trp Glu Ala Glu Glu Gly Gly Ser Pro Glu
                  260
                                       265
 Val Arg Thr Ser Leu Ala Asn Met Val Asn Pro Val Ser Thr Lys
                  275
                                      280
 Asn Thr Lys Ile Ser Gln Glu Leu Cys Ala Val Ile Pro Ala Thr
                  290
                                      295
 Trp Glu Ala Glu Val Gly Glu Leu Leu Glu Pro Gly Ser Trp Arg
                  305
                                      310
 Phe Gln
 <210> 156
 <211> 617
 <212> PRT
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 Ser Leu Lys Trp Gly Ser Gly Gly Arg Glu Thr Ala Ser Arg Gly
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Ala Trp Lys Val Val Lys Pro Glu Ser Asn Asp Lys Glu Thr Glu
                                                           15
                  20
                                       25
Ala Ala Tyr Glu Ser Asp Ile Pro Glu Glu Leu Cys Gly His His
                                      40
Leu Pro Gln Gln Ser Leu Lys Ser Tyr Asn Asp Ser Pro Asp Val
                                      55
Ile Val Glu Ala Gln Phe Asp Gly Ser Asp Ser Glu Asp Gly His
                  65
                                      70
Gly Ile Thr Gln Asn Val Leu Val Asp Gly Val Lys Lys Leu Ser
                  80
                                      85
Val Cys Val Ser Glu Lys Gly Arg Glu Asp Gly Asp Ala Pro Val
                  95
                                     100
Thr Lys Asp Glu Thr Thr Cys Ile Ser Gln Asp Thr Arg Ala Leu
                110
                                     115
Ser Glu Lys Ser Leu Gln Arg Ser Ala Lys Val Val Tyr Ile Leu
                125
                                     130
Glu Lys Lys His Ser Arg Ala Ala Thr Gly Phe Leu Lys Leu Leu
                140
                                     145
Ala Asp Lys Asn Ser Glu Leu Phe Arg Lys Tyr Ala Leu Phe Ser
                155
                                     160
Pro Ser Asp His Arg Val Pro Arg Ile Tyr Val Pro Leu Lys Asp
                170
                                     175
Cys Pro Gln Asp Phe Val Ala Arg Pro Lys Asp Tyr Ala Asn Thr
                185
                                    190
Leu Phe Ile Cys Arg Ile Val Asp Trp Lys Glu Asp Cys Asn Phe
```

				200					205	_	_			210
Ala	Leu	Gly	Gln	Leu 215	Ala	Lys	Ser	Leu	Gly 220	Gln	Ala	Gly	Glu	11e 225
Glu	Pro	Glu	Thr	Glu 230	Gly	Ile	Leu	Thr	Glu 235	Tyr	Gly	Val	Asp	Phe 240
Ser	Asp	Phe	Ser		Glu	Val	Leu	Glu		Leu	Pro	Gln	Gly	Leu 255
Pro	Trp	Thr	Ile		Pro	Glu	Glu	Phe		Lys	Arg	Arg	Asp	
Arg	Lys	Asp	Cys		Phe	Thr	Ile	Asp		Ser	Thr	Ala	Arg	
Leu	Asp	Asp	Ala		Ser	Cys	Lys	Pro		Ala	Asp	Gly	Asn	
Lys	Val	Gly	Val		Ile	Ala	Asp	Val		Tyr	Phe	Val	Pro	
Gly	Ser	Asp	Leu	Asp 320	Lys	Val	Ala	Ala		Arg	Ala	Thr	Ser	
Tyr	Leu	Val	Gln		Val	Val	Pro	Met		Pro	Arg	Leu	Leu	
Glu	Glu	Leu	Cys		Leu	Asn	Pro	Met		Asp	Lys	Leu	Thr	
Ser	Val	Ile	Trp		Leu	Thr	Pro	Glu		Lys	Ile	Leu	Asp	
Trp	Phe	Gly	Arg		Ile	Ile	Arg	Ser		Thr	Lys	Leu	Ser	Tyr 390
Glu	His	Ala	Gln	Ser 395	Met	Ile	Glu	Ser		Thr	Glu	Lys	Ile	
Ala	Lys	Glu	Leu		Pro	Ile	Ser	Pro		His	Ser	Ser	Glu	
Val	His	Gln	Ala	Val 425	Leu	Asn	Leu	His		Ile	Ala	Lys	Gln	
Arg	Gln	Gln	Arg			Asp	Gly	Ala		Arg	Leu	Asp	Gln	
Lys	Leu	Ala	Phe		Leu	Asp	His	Glu		Gly	Leu	Pro	Gln	
Сув	His	Ile	Tyr		Туг	Arg	Glu	Ser		Lys	Leu	Val	Glu	Glu 480
Phe	Met	Leu	Leu		Asn	Met	Ala	Val		His	Lys	Ile	His	Arg 495
Ala	Phe	Pro	Glu		Ala	Leu	Leu	Arg		His	Pro	Pro	Pro	Gln 510
Thr	Arg	Met	Leu		Asp	Leu	Val	Glu		Cys	Asp	Gln	Met	Gly 525
Leu	Pro	Val	Asp		Ser	Ser	Ala	. Gly		Leu	Asn	Lys	Ser	Leu 540
Thr	Gln	Thr	Phe		Asp	Asp	Lys	Туг		Leu	Ala	Arg	Lys	Glu 555
Val	Leu	Thr	Asr		Суя	Ser	Arg	Pro		Gln	Met	Ala	Leu	Tyr 570
Phe	е Сув	Ser	Gly		ı Leı	ı Gln	Asp	Pro		Gln	Phe	Arg	, His	Tyr 585
Ala	. Lev	. Asn	val		Sei	. Val	His	Thr		His	Lev	ı Ala	His	Pro 600
Pro	Lev	ı Cys	Arg		y Pro	Gly	Ala	Pro		Pro	G13	у Сув	arg	Val 615
Arg	J Lev	ı		500	-									

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<212> PRT

<213> Homo sapiens

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Tyr Leu Asn Leu Leu Val Thr Ser Trp Arg Met Asn Asp Ser Leu
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                  5
  1
Val Ile Gln Gln Asn Asp Leu Val Phe Glu Phe Ala Ser Asn Val
                 20
Met Glu Asp Glu Arg Gln Leu Gly Asp Pro Ala Ile Phe Pro Ala
                                      40
Val Ile Val Glu His Val Pro Gly Ala Asp Ile Leu Asn Ser Tyr
                                      55
                 50
Ala Gly Leu Ala Cys Val Glu Glu Pro Asn Asp Met Ile Thr Glu
                                      70
                 65
Ser Ser Leu Asp Val Ala Glu Glu Glu Ile Ile Asp Asp Asp Asp
                 80
                                      85
Asp Asp Ile Thr Leu Thr Val Glu Ala Ser Cys His Asp Gly Asp
                 95
                                     100
Glu Thr Ile Glu Thr Ile Glu Ala Ala Glu Ala Leu Leu Asn Met
                                     115
                                                         120
                110
Asp Ser Pro Gly Pro Met Leu Asp Glu Lys Arg Ile Asn Asn Asn
                125
                                     130
Ile Phe Ser Ser Pro Glu Asp Asp Met Val Val Ala Pro Val Thr
                140
                                     145
His Val Ser Val Thr Leu Asp Gly Ile Pro Glu Val Met Glu Thr
                155
                                     160
Gln Gln Val Gln Glu Lys Tyr Ala Asp Ser Pro Gly Ala Ser Ser
                                                          180
                170
                                     175
Pro Glu Gln Pro Lys Arg Lys Lys Gly Arg Lys Thr Lys Pro Pro
                                     190
                185
Arg Pro Asp Ser Pro Ala Thr Thr Pro Asn Ile Ser Val Lys
                                     205
                 200
Lys Asn Lys Asp Gly Lys Gly Asn Thr Ile Tyr Leu Trp Glu Phe
                215
                                     220
Leu Leu Ala Leu Leu Gln Asp Lys Ala Thr Cys Pro Lys Tyr Ile
                 230
                                     235
Lys Trp Thr Gln Arg Glu Lys Gly Ile Phe Lys Leu Val Asp Ser
                245
                                     250
                                                          255
Lys Ala Val Ser Arg Leu Trp Gly Lys His Lys Asn Lys Pro Asp
                 260
                                     265
                                                          270
Met Asn Tyr Glu Thr Met Gly Arg Ala Leu Arg Tyr Tyr Gln
                                                          285
                 275
                                     280
Arg Gly Ile Leu Ala Lys Val Glu Gly Gln Arg Leu Val Tyr Gln
                 290
                                     295
Phe Lys Glu Met Pro Lys Asp Leu Ile Tyr Ile Asn Asp Glu Asp
                 305
                                     310
                                                          315
Pro Ser Ser Ser Ile Glu Ser Ser Asp Pro Ser Leu Ser Ser Ser
                 320
                                     325
Ala Thr Ser Asn Arg Asn Gln Thr Ser Arg Ser Arg Val Ser Ser
                 335
                                     340
                                                          345
Ser Pro Gly Val Lys Gly Gly Ala Thr Thr Val Leu Lys Pro Gly
                 350
                                     355
Asn Ser Lys Ser Cys Lys Ser Gln Arg Ser Cys
                 365
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<211> 871
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<213> Homo sapiens
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<223> Incyte ID No: LG:257088.20.orf2:2002JAN18

<400> 158 Arg Leu Phe Val Leu Ile Ser Leu Glu Leu Lys Met Leu Tyr Phe Ser Arg Ser His Phe Pro Arg Pro Cys Gly Gly Gln Val Ser Ala Gly Ser Gly Leu Thr Leu Leu Leu Leu Leu Pro Ala Leu Trp Arg Gly Trp Leu Glu Gly Asp Gly Gln Gln Ala Val Pro Ala Arg Gly Glu Pro Gln Gln Asp Cys Cys Val Lys Thr Glu Leu Leu Gly Glu Glu Thr Pro Met Ala Ala Asp Glu Gly Ser Ala Glu Lys Gln Ala Gly Glu Ala His Met Ala Ala Asp Gly Glu Thr Asn Gly Ser Cys Glu Asn Ser Asp Ala Ser Ser His Ala Asn Ala Ala Lys His Thr Gln Asp Ser Ala Arg Val Asn Pro Gln Asp Gly Thr Asn Thr Leu Thr Arg Ile Ala Glu Asn Gly Val Ser Glu Arg Asp Ser Glu Ala Ala Lys Gln Asn His Val Thr Ala Asp Asp Phe Val Gln Thr Ser Val Ile Gly Ser Asn Gly Tyr Ile Leu Asn Lys Pro Ala Leu Gln Ala Gln Pro Leu Arg Thr Thr Ser Thr Leu Ala Ser Ser Leu Pro Gly His Ala Ala Lys Thr Leu Pro Gly Gly Ala Gly Lys Gly Arg Thr Pro Ser Ala Phe Pro Gln Thr Pro Ala Ala Pro Pro Ala Thr Leu Gly Glu Gly Ser Ala Asp Thr Glu Asp Arg Lys Leu Pro ·235 Ala Pro Gly Ala Asp Val Lys Val His Arg Ala Arg Lys Thr Met Pro Lys Ser Val Val Gly Leu His Ala Ala Ser Lys Asp Pro Arg Glu Val Arg Glu Ala Arg Asp His Lys Glu Pro Lys Glu Glu Ile Asn Lys Asn Ile Ser Asp Phe Gly Arg Gln Gln Leu Leu Pro Pro Phe Pro Ser Leu His Gln Ser Leu Pro Gln Asn Gln Cys Tyr Met Ala Thr Thr Lys Ser Gln Thr Ala Cys Leu Pro Phe Val Leu Ala Ala Ala Val Ser Arg Lys Lys Arg Arg Met Gly Thr Tyr Ser Leu Val Pro Lys Lys Lys Thr Lys Val Leu Lys Gln Arg Thr Val Ile Glu Met Phe Lys Ser Ile Thr His Ser Thr Val Gly Ser Lys Gly Glu Lys Asp Leu Gly Ala Ser Ser Leu His Val Asn Gly Glu Ser Leu Glu Met Asp Ser Asp Glu Asp Asp Ser Glu Glu Leu Glu Glu Asp Asp Gly His Gly Ala Glu Gln Ala Ala Ala Phe Pro Thr Glu Asp Ser Arg Thr Ser Lys Glu Ser Met Ser Glu Ala Asp Arg Ala Gln Lys Ser Ser Glu Ser Ser Ile Lys Lys Lys Phe Leu Lys 

Arg	Lys	Gly	Lys	Thr 455	Asp	Ser	Pro	Trp	Ile 460		Pro	Ala	Arg	Lys 465
Arg	Arg	Arg	Arg	Ser 470	Arg	Lys	Lys	Pro	Ser	Gly	Ala	Leu	Gly	Ser 480
Glu	Ser	Tyr	Lys	Ser 485	Ser	Ala	Gly	Ser		Glu	Gln	Thr	Ala	Pro 495
Gly	Asp	Ser	Thr		Tyr	Met	Glu	Val		Leu	Asp	Ser	Leu	Asp 510
Leu	Arg	Val	Lys		Ile	Leu	Ser	Ser		Ala	Glu	Gly	Leu	Ala 525
Asn	Gly	Pro	Asp		Leu	Glu	Thr	Asp		Leu	Gln	Glu	Val	Pro 540
Leu	Cys	Ser	Суѕ		Met	Glu	Thr	Pro		Ser	Arg	Glu	Ile	Thr 555
Thr	Leu	Ala	Asn	Asn 560	Gln	Cys	Met	Ala		Glu	Ser	Val	Asp	His 570
Glu	Gly	Asn	Phe		Glu	Cys	Gln	Pro		Ser	Ser	Ile	Ser	His 585
Arg	Phe	His	Lys	Asp 590	Суѕ	Ala	Ser	Arg		Asn	Asn	Ala	Ser	Tyr 600
Cys	Pro	His	Суз	Gly 605	Glu	Glu	Ser	Ser		Ala	Lys	Glu	Va1	Thr 615
	Ala			620					625					Gly 630
	Glu			635					640					Thr 645
	Ser		•	650			. 1		655					Gln 660
	Ala			665					670				_	Pro 675
	Gly			680					685			•		Gly 690
	Glu			695					700			•		705
	Lys			710	;				715					720
	Gln			725					730					Gly 735
	Asp			740					745					Pro 750
	His			755					760					765
	Val			770					775					780
	Thr			785					790					795
	Lys			800					805					810
	Glu			815					820					825
	Glu			830					835					840
	Cys			845					850				-	855
	Val	GIII	ATS	860	чтλ	Pro	Arg	GLu	Ala 865	Ala	Ala	Val	Gln	Gly 870
Leu														

<210> 159 <211> 157 <212> PRT <213> Homo sapiens

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Thr Ile Ala Tyr Leu Leu Ile Lys Ser Lys Cys Leu Ser Leu Ala
Val Pro Pro Leu Leu Ser Gly Asn Asp Phe Gln Thr Val Glu Glu
Gly Ser Asn Val Lys Leu Val Cys Asn Val Lys Ala Asn Pro Gln
                                     40
                 35
Ala Gln Met Met Trp Tyr Lys Asn Ser Ser Leu Leu Asp Leu Glu
                                     55
                 50
Lys Ser Arg His Gln Ile Gln Gln Thr Ser Glu Ser Phe Gln Leu
                                     70
                 65
Ser Ile Thr Lys Val Glu Lys Pro Asp Asn Gly Thr Tyr Ser Cys
                                     85
                 80
Ile Ala Lys Ser Ser Leu Lys Thr Glu Ser Leu Asp Phe His Leu
                                     100
                 95
Ile Val Lys Asp Lys Thr Val Gly Val Pro Ile Glu Pro Ile Ile
                                    115
                110
Ala Ala Cys Val Val Ile Phe Leu Thr Leu Cys Phe Gly Leu Ile
                                    130
                125
Ala Arg Arg Lys Lys Ile Met Lys Leu Cys Met Lys Asp Lys Asp
                140
                                    145
Pro His Ser Glu Thr Ala Leu
                155
<210> 160
<211> 280
<212> PRT
<213> Homo sapiens
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<223> Incyte ID No: LG:275355.12.orf1:2002JAN18
<400> 160
Lys Tyr Glu Phe Asp Asp Tyr Glu Arg Phe Ile Lys Tyr Leu Gly
                                      10
Gly Leu Asn Phe Met Thr Thr Leu Thr Thr His Lys His Leu Pro
                 20
                                      25
His Arg Arg Val Ser Pro Asp Leu Leu Ile Leu Pro Cys Thr Phe
                                      40
                 35
Ala Ser Val Gly Ile Met Trp Ile Asp Ser Val Phe Phe Arg Leu
                                      55
                 50
Val Asp Ala Leu Lys Leu Gln Asp Gln Leu Lys Ala Pro Val Lys
                                                          75
                                      70
                 65
Thr Leu Ser Glu Gly Ile Lys Arg Lys Leu Cys Phe Val Leu Ser
                                      85
                 80
Ile Leu Gly Asn Pro Ser Val Val Leu Leu Asp Glu Pro Ser Thr
                                     100
                  95
Gly Met Asp Pro Glu Gly Gln Gln Met Trp Gln Val Ile Arg
                                     115
                 110
Ala Thr Phe Arg Asn Thr Glu Arg Gly Ala Leu Leu Thr Thr His
                 125
                                     130
Tyr Met Ala Glu Ala Glu Ala Val Cys Asp Arg Val Ala Ile Met
                                                          150
                                     1.45
                 140
Val Ser Gly Arg Leu Arg Cys Ile Gly Ser Ile Gln His Leu Lys
                                     160
                 155
 Ser Lys Phe Gly Lys Asp Tyr Leu Leu Glu Met Lys Leu Lys Asn
                                     175
                 170
Leu Ala Gln Met Glu Pro Leu His Ala Glu Ile Leu Arg Leu Phe
```

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185
                                     190
                                                         195
Pro Gln Ala Ala Gln Gln Glu Arg Phe Ser Ser Leu Met Val Tyr
                 200
                                     205
                                                         210
Lys Leu Pro Val Glu Asp Val Arg Pro Leu Ser Gln Ala Phe Phe
                 215
                                     220
Lys Leu Glu Ile Val Lys Gln Ser Phe Asp Leu Glu Glu Tyr Ser
                 230
                                     235
Leu Ser Gln Ser Thr Leu Glu Gln Val Phe Leu Glu Leu Ser Lys
                 245
                                     250
Glu Gln Glu Leu Gly Asp Leu Glu Glu Asp Phe Asp Pro Ser Val
                 260
                                     265
Lys Trp Lys Leu Leu Gln Glu Glu Pro
                275
<210> 161
<211> 149
<212> PRT
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Met Trp Ile Val Asp Ser Asn Ile Ile Thr Ala Ile Val Gln Leu
  1
                   5
                                      10
His Gly Leu Trp Met Asp Cys Thr Trp Tyr Ser Thr Gly Met Phe
                 20
                                      25
Ser Cys Ala Leu Lys His Ser Ile Leu Ser Leu Pro Ile His Val
                 35
                                      40
Gln Ala Ala Arg Ala Thr Met Val Leu Ala Cys Val Leu Ser Ala
                 50
Leu Gly Ile Cys Thr Ser Thr Val Gly Met Lys Cys Thr Arg Leu
                 65
                                      70
Gly Gly Asp Arg Glu Thr Lys Ser His Ala Ser Phe Ala Gly Gly
                 80
                                      85
                                                          90
Val Cys Phe Met Ser Ala Gly Ile Ser Ser Leu Ile Ser Thr Val
                 95
                                     100
Trp Tyr Thr Lys Glu Ile Ile Ala Asn Phe Leu Asp Leu Thr Val
                110
                                     115
Pro Glu Ser Asn Lys His Glu Pro Gly Gly Ala Ile Tyr Ile Gly
                125
                                     130
Phe Ile Ser Ala Met Leu Leu Phe Ile Ser Gly Met Ile Phe
                140
                                     145
<210> 162
<211> 281
<212> PRT
<213> Homo sapiens
<220>
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<223> Incyte ID No: LG:299937.3.orf3:2002JAN18
<400> 162
Phe Gly Gly Arg Pro Ala Gly Ala Ser Pro Leu Leu Ser Ser Lys
                                      10
Leu Thr Tyr Leu His Leu Pro Ala Gly Ile Lys Met Ala Gly Tyr
                 20
                                      25
Ala Thr Thr Pro Ser Pro Met Gln Thr Leu Gln Glu Glu Ala Val
                 35
                                     40
Cys Ala Ile Cys Leu Asp Tyr Phe Lys Asp Pro Val Ser Ile Ser
                                     55
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Cys Gly His Asn Phe Cys Arg Gly Cys Val Thr Gln Leu Trp Ser
                 65
Lys Glu Asp Glu Glu Asp Gln Asn Glu Glu Glu Asp Glu Trp Glu
                                      85
                 80
Glu Glu Glu Asp Glu Glu Ala Val Gly Ala Met Asp Gly Trp Asp
                                     100
                 95
Gly Ser Ile Arg Glu Val Leu Tyr Arg Gly Asn Ala Asp Glu Glu
                                                         120
                                     115
                110
Leu Phe Gln Asp Gln Asp Asp Glu Leu Trp Leu Gly Asp Ser
                                     130
                125
Gly Ile Thr Asn Trp Asp Asn Val Asp Tyr Met Trp Asp Glu Glu
                                     145
Glu Glu Glu Glu Asp Gln Asp Tyr Tyr Leu Gly Gly Leu Arg
                                                         165
                                     160
                155
Pro Asp Leu Arg Ile Asp Val Tyr Arg Glu Glu Glu Ile Leu Glu
                170
                                     175
                                                         180
Ala Tyr Asp Glu Asp Glu Asp Glu Glu Leu Tyr Pro Asp Ile His
                                                          195
                                     190
                185
Pro Pro Pro Ser Leu Pro Leu Pro Gly Gln Phe Thr Cys Pro Gln
                                     205
                                                          210
                200
Cys Arg Lys Ser Phe Thr Arg Arg Ser Phe Arg Pro Asn Leu Gln
                                     220
                                                          225
                215
Leu Ala Asn Met Val Gln Ile Ile Arg Gln Met Cys Pro Thr Pro
                                     235
                230
Tyr Arg Gly Asn Arg Ser Asn Asp Gln Gly Met Cys Phe Lys His
                                     250
                245
Gln Glu Ala Leu Lys Leu Phe Cys Glu Val Asp Lys Glu Ala Ile
                 260
                                     265
Cys Val Val Cys Arg Glu Ser Arg Ser His Lys
                275
<210> 163
<211> 703
<212> PRT
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 Gly Arg Ser Ser Pro Arg Ala Arg Leu Arg Gly Trp Thr Leu Arg
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 Ala Pro Gly Lys Glu Thr Pro Ala Phe Ala Thr Met Leu Ser Ser
                                       25
                  20
 Thr Asp Phe Thr Phe Ala Ser Trp Glu Leu Val Val Arg Val Asp
                                                           45
                                       40
                  35
 His Pro Asn Glu Glu Gln Gln Lys Asp Val Thr Leu Arg Val Ser
                                       55
                  50
 Gly Asp Leu His Val Gly Gly Val Met Leu Lys Leu Val Glu Gln
                  · 65
 Ile Asn Ile Ser Gln Asp Trp Ser Asp Phe Ala Leu Trp Trp Glu
                                       85
 Gln Lys His Cys Trp Leu Leu Lys Thr His Trp Thr Leu Asp Lys
                                      100
                  95
 Tyr Gly Val Gln Ala Asp Ala Lys Leu Leu Phe Thr Pro Gln His
                 110
                                      115
 Lys Met Leu Arg Leu Arg Leu Pro Asn Leu Lys Met Val Arg Leu
                                      130
                 125
 Arg Val Ser Phe Ser Ala Val Val Phe Lys Ala Val Ser Asp Ile
                                                           150
                                      145
                 140
 Cys Lys Ile Leu Asn Ile Arg Arg Ser Glu Glu Leu Ser Leu Leu
                 155
                                      160
```

Lys	s Pro	Se:	r Gl	y Asp 170	э Тул Э	Phe	∋ Lys	5 Lys	5 Lys 175	Lys	Lys	Lys	s Ası	Lys
				185	)				11e	Leu				180 Ser 195
				200	)				205	,				Ser
				215	)				11e	Asr				Ala 225
				230	}				235	1				Gln
				245	)				250	+				Glu 255
				260	l.				265					233 Ala 270
				275					280					Glu 285
				290					295					Tyr 300
				305					310					Ile 315
				320					325					Glu 330
				Glu 335					340					3/5
				Lys 350					355					360
				Glu 365					370					Asn
				Leu 380					385					Glu
				Ile 395					400					Phe
				Leu 410					415					Phe
				Thr 425					430					Leu 435
				Pro 440					Asn 445					Glu
				Val 455					460					Lys
				Val 470					475					Arg
				Asn 485					490					Met
				Gly 500					505					Pro
				Ile 515					Arg 520					Asn
				Val 530					Glu 535					Asn
				Val 545					Ala 550					Ser
				Ala 560					Ala 565					Ala
Gln	Met	Pro	Leu	Val 575	Glu	Ala	Lys	Leu	Arg 580	Phe	Ile	Gln	Ala	Trp
Gln	Ser	Leu	Pro	Glu 590	Phe	Gly	Leu	Thr	Tyr 595	Tyr	Leu	Val	Arg	
Lys	Gly	Ser	Lys	Lys 605	Asp	Asp	Ile	Leu	Gly 610	Val	Ser	Tyr	Asn	
Leu	Ile	Lys	Ile		Ala	Ala	Thr	Gly	Ile 625	Pro	Val	Thr	Thr	
Arg	Phe	Thr	Asn		Lys	Gln	Trp	Asn	Val	Asn	Trp	Glu	Thr	630 Arg

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640
                635
Gln Val Val Ile Glu Phe Asp Gln Asn Val Phe Thr Ala Phe Thr
                                    655
                650
Cys Leu Ser Ala Asp Cys Lys Ile Val His Glu Tyr Ile Gly Gly
                665
                                    670
Tyr Ile Phe Leu Ser Thr Arg Ser Lys Asp Gln Asn Glu Thr Leu
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                680
Asp Glu Asp Leu Phe His Lys Leu Thr Gly Gly Gln Asp
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Gly Thr His Pro Ser Thr Val Leu Leu Ser Pro Leu Ala Gly Val
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Glu Leu Pro Val Tyr Asp Ile Thr Lys Lys His Leu Ile Leu Ser
                                      25
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Gly Leu Met Gly Asp Thr Val Tyr Thr His Phe Leu Ser Ser Phe
                                      40
                 35
Thr Cys Gly Leu Ala Gly Ala Leu Ala Ser Asn Pro Val Asp Val
                 50
                                      55
Val Arg Thr Arg Met Met Asn Gln Arg Val Leu Arg Asp Gly Arg
                 65
Cys Ser Gly Tyr Thr Gly Thr Leu Asp Cys Leu Leu Gln Thr Trp
                                      85
Lys Asn Glu Gly Phe Phe Ala Leu Tyr Lys Gly Phe Trp Pro Asn
                 95
                                     100
Trp Leu Arg Leu Gly Pro Trp Asn Ile Ile Phe Phe Val Thr Tyr
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Glu Gln Leu Lys Lys Leu Asp Leu
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Val Asp Glu His Leu His Glu Asp Asn Gly Glu Val Glu Val Arg
Arg Ser Cys Arg Ile Arg Ser Arg Tyr Ser Gly Val Asn Gln Ser
Met Leu Phe Asp Lys Leu Ile Thr Asn Thr Ala Glu Ala Val Leu
                  35
                                      40
Gln Lys Met Asp Asp Met Lys Lys Met Arg Arg Gln Arg Met Arg
                                      55
Glu Leu Glu Asp Leu Gly Val Phe Asn Glu Thr Glu Glu Ser Asn
                                      70
                  65
Leu Asn Met Tyr Thr Arg Gly Lys Gln Lys Asp Ile Gln Arg Thr
                                      85
                  80
Asp Glu Glu Thr Thr Asp Asn Gln Glu Gly Ser Val Glu Ser Ser
                  95
                                     100
                                                          105
```

Glu Glu Gly Glu Asp Gln Glu His Glu Asp Asp Gly Glu Asp Glu Asp Asp Glu Asp Glu Asp Glu Glu Asp Gly Glu Glu Asn Gln Lys Arg Tyr Tyr Leu Arg Gln Arg Lys Ala Thr Val Tyr Tyr Gln Ala Pro Leu Glu Lys Pro Arg His Gln Arg Lys Pro Asn Ile Phe Tyr Ser Gly Pro Ala Ser Pro Ala Arg Pro Arg Tyr Arg Leu Ser Ser Ala Gly Pro Arg Ser Pro Tyr Cys Lys Arg Met Asn Arg Arg Arg His Ala Ile His Ser Ser Asp Ser Thr Ser Ser Ser Ser Glu Asp Glu Gln His Phe Glu Arg Arg Arg Lys Arg Ser Arg Asn Arg Ala Ile Asn Arg Cys Leu Pro Leu Asn Phe Arg Lys Asp Glu Leu Lys Gly Ile Tyr Lys Asp Arg Met Lys Ile Gly Ala Ser Leu Ala Asp Val Asp Pro Met Gln Leu Asp Ser Ser Val Arg Phe Asp Ser Val Gly Gly Leu Ser Asn His Ile Ala Ala Leu Lys Glu Met Val Val Phe Pro Leu Leu Tyr Pro Glu Val Phe Glu Lys Phe Lys Ile Gln Pro Pro Arg Gly Cys Leu Phe Tyr Gly Pro Pro Gly Thr Gly Lys Thr Leu Val Ala Arg Ala Leu Ala Asn Glu Cys Ser Gln Gly Asp Lys Arg Val Ala Phe Phe Met Arg Lys Gly Ala Asp Cys Leu Ser Lys Trp Val Gly Glu Ser Glu Arg Gln Leu Arg Leu Leu Phe Asp Gln Ala Tyr Gln Met Arg Pro Ser Ile Ile Phe Phe Asp Glu Ile Asp Gly Leu Ala Pro Val Arg Ser Ser Arg Gln Asp Gln Ile His Ser Ser Ile Val Ser Thr Leu Leu Ala Leu Met Asp Gly Leu Asp Ser Arg Gly Glu Ile Val Val Ile Gly Ala Thr Asn Arg Leu Asp Ser Ile Asp Pro Ala Leu Arg Arg Pro Gly Arg Phe Asp Arg Glu Phe Leu Phe Ser Leu Pro Asp Lys Glu Ala Arg Lys Glu Ile Leu Lys Ile His Thr Arg Asp Trp Asn Pro Lys Pro Leu Asp Thr Phe Leu Glu Glu Leu Ala Glu Asn Cys Val Gly Tyr Cys Gly Ala Asp Ile Lys Ser Ile Cys Ala Glu Ala Ala Leu Cys Ala Leu Arg Arg Arg Tyr Pro Gln Ile Tyr Thr Thr Ser Glu Lys Leu Gln Leu Asp Leu Ser Ser Ile Asn Ile Ser Ala Lys Asp Phe Glu Val Ala Met Gln Lys Met Ile Pro Ala Ser Gln Arg Ala Val Thr Ser Pro Gly Gln Ala Leu Ser Thr Val Val Lys Pro Leu Leu Gln Asn Thr Val Asp Lys Ile Leu Glu Ala Leu Gln Arg Val Phe Pro PCT/US03/01363 WO 03/062379

580 585
Jis Ala Clu Phe Arg Thr Asn Lys Thr Leu Asp Ser Asp Ile Ser
590 Cor Asp Leu Ala Tyr Ser Asp Asp Val
Cys Pro Leu Leu Glu Ser Asp Bed 1610 615 610 615
605 Pro Ser Val Tyr Glu Asn Gly Leu Ser Gln Lys Ser Ser His Lys 630 625 627
Ala Lys Asp Asn Phe Asn Phe Leu His Leu Asn Arg Asn Ala Cys 640 645
mm Cla Pro Met Ser Phe Arg Pro Arg Ile Leu Ile Val Gly Glu
650  Pro Gly Phe Gly Gln Gly Ser His Leu Ala Pro Ala Val Ile His 670  675
675 675 676 677 678 From The Lett Asp Tle Pro Val Lett
665 Ala Leu Glu Lys Phe Thr Val Tyr Thr Leu Asp Ile Pro Val Leu 690 680 685
680 Phe Gly Val Ser Thr Thr Ser Pro Glu Glu Thr Cys Ala Gln Val 705 695 700 870 870 870 870 870
Ile Arg Glu Ala Lys Arg Thr Ala Pro Ser Ile Val Tyl Val Flo
The His Val Trp Trp Glu Ile Val Gly Pro Thr Leu Lys Ala
730 735 725 725 730 730 Thr Phe Thr Thr Leu Gln Asn Ile Pro Ser Phe Ala Pro Val 745 750
Thr Phe Thr Thr Hed Hed Sun 745  740  745  740  750  745  750  750
740  Leu Leu Leu Ala Thr Ser Asp Lys Pro His Ser Ala Leu Pro Glu 765 755  760 760 760 760 760 760 760 760 760 76
755 Glu Val Gln Glu Leu Phe Ile Arg Asp Tyr Gly Glu Ile Phe Asn 780
770 775 Val Gln Leu Pro Asp Lys Glu Glu Arg Thr Lys Phe Phe Glu Asp 795 795
Leu Ile Leu Lys Gln Ala Ala Lys Pro Pro Ile Ser Lys Lys 810
800 805 Ala Val Leu Gln Ala Leu Glu Val Leu Pro Val Ala Pro Pro 825
815 Glu Pro Arg Ser Leu Thr Ala Glu Glu Val Lys Arg Leu Glu Glu 835 840
Gla Clu Clu Asp Thr Phe Arg Glu Leu Arg Ile Phe Leu Arg Asn
850 845 845 Val Thr His Arg Leu Ala Ile Asp Lys Arg Phe Arg Val Phe Thr 865 870
Val Thr His Arg Bed Ard 125 865 870 865 870 860 Rep Tyr Val Thr Val Ile
860 Lys Pro Val Asp Pro Asp Glu Val Pro Asp Tyr Val Thr Val Ile 885 875 880 880
Lys Gln Pro Met Asp Leu Ser Ser Val Ile Ser Lys Ile Asp Leu 900
890  His Lys Tyr Leu Thr Val Lys Asp Tyr Leu Arg Asp Ile Asp Leu 910 915
The Cys Ser Asn Ala Leu Glu Tyr Asn Pro Asp Arg Asp Pro Gly
920 920 925 Asp Arg Leu Ile Arg His Arg Ala Cys Ala Leu Arg Asp Thr Ala 945
Asp Arg Leu He Arg his 123 940 945 945 935 940 Sin Asp Phe Glu Gln Leu
935  Tyr Ala Ile Ile Lys Glu Glu Leu Asp Glu Asp Phe Glu Gln Leu 960 950  955  957  958  958  958  958
950 950 Cys Glu Glu Ile Gln Glu Ser Arg Lys Lys Arg Gly Cys Ser Ser 975 965 970 970 970 970 970
965  Ser Lys Tyr Ala Pro Ser Tyr Tyr His Val Met Pro Lys Gln Asn 990 980 985
Ser Thr Leu Val Gly Asp Lys Arg Ser Asp Pro Glu GIN ASN GIV
Lys Leu Lys Thr Pro Ser Thr Pro Val Ala Cys Ser Thr F10 1120
Gln Leu Lys Arg Lys Ile Arg Lys Lys Ser Ash Tip Tyl Led 1035
1025 1030 Thr Ile Lys Lys Arg Arg Lys Ile Ser Gln Ala Lys Asp Asp Ser 1050
1040

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Gln Asn AÌa Ile Asp His Lys Ile Glu Ser Asp Thr Glu Glu Thr
                 1055
                                      1060
  Gln Asp Thr Ser Val Asp His Asn Glu Thr Gly Asn Thr Gly Glu
                 1070
                                     1075
  Ser Ser Val Glu Glu Asn Glu Lys Gln Gln Asn Ala Ser Glu Ser
                 1085
                                     1090
  Lys Leu Glu Leu Arg Asn Asn Ser Asn Thr Cys Asn Ile Glu Asn
                 1100
                                     1105
  Glu Leu Glu Asp Ser Arg Lys Thr Thr Ala Cys Thr Glu Leu Arg
                 1115
                                     1120
  Asp Lys Ile Ala Cys Asn Gly Asp Ala Ser Ser Ser Gln Ile Ile
                 1130
                                     1135
 His Ile Ser Asp Glu Asn Glu Gly Lys Glu Met Cys Val Leu Arg
                 1145
                                     1150
 Met Thr Arg Ala Arg Arg Ser Gln Val Glu Gln Gln Leu Ile
                                                         1155
                1160
                                     1165
 Thr Val Glu Lys Ala Leu Ala Ile Leu Ser Gln Pro Thr Pro Ser
                1175
                                     1180
 Leu Val Val Asp His Glu Arg Leu Lys Asn Leu Leu Lys Thr Val
                1190
                                     1195
 Val Lys Lys Ser Gln Asn Tyr Asn Ile Phe Gln Leu Glu Asn Leu
                1205
                                     1210
 Tyr Ala Val Ile Ser Gln Cys Ile Tyr Arg His Arg Lys Asp His
                1220
                                     1225
 Asp Lys Thr Ser Leu Ile Gln Lys Met Glu Gln Glu Val Glu Asn
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 Phe Ser Cys Ser Arg
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His Met Val Ile His Thr Gly Leu Lys Ser His Gln Cys Pro Leu
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Cys Pro Phe Arg Cys Ala Arg Lys Asp Asn Leu Lys Ser His Met
                  35
Lys Val His Gln His Gln Asp Arg Gly Glu Thr Phe Gln Cys Gln
                 50
                                      55
Leu Cys Pro Phe Thr Ser Ser Arg His Phe Ser Leu Lys Leu His
                 65
                                      70
Met Arg Cys His Gln His Phe Leu Arg Thr Glu Ala Lys Val Lys
                 80
                                      85
Glu Glu Ile Pro Asp Pro Asp Val Lys Gly Ser Pro His Leu Ser
                 95
                                     100
Asp Ser Ala Cys Leu Gly Gln Gln Arg Glu Gly Gly Thr Glu
                110
                                     115
Leu Val Gly Thr Met Met Thr Ser Asn Thr Pro Glu Arg Thr Ser
                125
                                    130
Gln Gly Gly Ala Gly Val Ser Pro Leu Leu Val Lys Glu Glu Pro
                140
                                    145
Lys Glu Asp Asn Gly Leu Pro Thr Ser Phe Thr Leu Asn Ala Ala
                155
                                    160
Asp Arg Pro Ala Asn His Thr Lys Leu Lys Asp Pro Ser Glu Tyr
                170
                                    175
                                                         180
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Val	Ala	Asn	Ser	Ala 185	Ser	Ala	Leu	Phe	Ser 190	Gln	Asp	Ile	Ser	Val 195
Lys	Met	Ala	Ser		Phe	Leu	Met	Lys	Leu 205	Ser	Ala	Ala	Asn	Gln 210
Lys	Glu	Pro	Met		Leu	Asn	Phe	Lys	Val 220	Lys	Glu	Glu	Pro	Lys 225
Glu	Gly	Glu	Ser		Ser	Thr	Thr	Leu	Pro 235	Arg	Ser	Ser	Tyr	Val 240
Phe	Ser	Pro	Glu	Ser 245		Val	Ser	Ala	Pro 250	Gly	Val	Ser	Glu	Asp 255
Ala	Leu	Lys	Pro	Gln 260	Glu	Gly	Lys	Gly	Ser 265	Val	Leu	Arg	Arg	Asp 270
				275			Glu		280					285
		_	_	290			Met		295					300
				305			Ser		310					315
_				320			Ļeu		325					330
				335			Gly		340					345
				350			Val		355					360
				365			Ser		370					375
				380			Thr		385					390
				395			Thr		400					405
				410			Gly	•	415					420
				425			Gly		430					435
				440			Сув		445					450
				455			Arg		460					465
				470			Leu		475					480
				485			Leu		490					495 Glu
				500					505					510 Val
				515					520					525 Met
				530					535					540 Cys
				545					550					555 Thr
				560	1				565					570 Ser
				575					580					585 Asp
				590	)				595					600 Lys
				605	,				610	1				615 Gln
				620	<b>)</b> .				625	,				630 Cys
				635	i				640	)				645 Arg
		-				_	_							

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650
                                     655
His Leu Gln Ile His Leu Ile Thr Arg Met Phe Glu Cys Asp Val
                                     670
Cys His Lys Phe Met Lys Thr Pro Glu Gln Leu Leu Glu His Lys
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                                     685
                                                          690
Lys Cys His Thr Val Pro Thr Gly Gly Leu Asn Ser Gly Gln Trp
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                                     700
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Ala Ala Pro Pro Ser Arg Arg His Ser Val Thr Phe Val Pro
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                                      25
Ser Gly Ala Ala Arg Gly Leu Ser Arg Met Val Pro Ser Ser Pro
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                                      40
Ala Val Glu Lys Gln Val Pro Val Glu Pro Gly Pro Asp Pro Glu
                  50
                                      55
Leu Arg Ser Trp Arg Arg Leu Val Cys Tyr Leu Cys Phe Tyr Gly
                  65
                                      70
                                                          75
Phe Met Ala Gln Ile Arg Pro Gly Glu Ser Phe Ile Thr Pro Tyr
                  80
                                                          90
Leu Leu Gly Pro Asp Lys Asn Phe Thr Arg Glu Gln Val Thr Asn
                  95
                                     100
Glu Ile Thr Pro Val Leu Ser Tyr Ser Tyr Leu Ala Val Leu Val
                110
                                     115
Pro Val Phe Leu Leu Thr Asp Tyr Leu Arg Tyr Thr Pro Val Leu
                125
                                     130
Leu Leu Gln Gly Leu Ser Phe Val Ser Val Trp Leu Leu Leu
                140
                                     145
Leu Gly His Ser Val Ala His Met Gln Leu Met Glu Leu Phe Tyr
                155
                                     160
                                                         165
Ser Val Thr Met Ala Ala Arg Ile Ala Tyr Ser Ser Tyr Ile Phe
                170
                                     175
Ser Leu Val Arg Pro Ala Arg Tyr Gln Arg Val Ala Gly Tyr Ser
                185
                                     190
                                                         195
Arg Ala Ala Val Leu Leu Gly Val Phe Thr Ser Ser Val Leu Gly
                200
                                     205
Gln Leu Leu Val Thr Val Gly Arg Val Ser Phe Ser Thr Leu Asn
                215
                                     220
Tyr Ile Ser Leu Ala Phe Leu Thr Phe Ser Val Val Leu Ala Leu
                230
                                     235
Phe Leu Lys Arg Pro Lys Arg Ser Leu Phe Phe Asn Arg Asp Asp
                245
                                     250
Arg Gly Arg Cys Glu Thr Ser Ala Ser Glu Leu Glu Arg Met Asn
                260
                                     265
Pro Gly Pro Gly Gly Lys Leu Gly His Ala Leu Arg Val Ala Cys
                275
                                    280
Gly Asp Ser Val Leu Ala Arg Met Leu Arg Glu Leu Gly Asp Ser
                290
                                     295
Leu Arg Arg Pro Gln Leu Arg Leu Trp Ser Leu Trp Trp Val Phe
                305
                                     310
Asn Ser Ala Gly Tyr Tyr Leu Val Val Tyr Tyr Val His Ile Leu
                320
                                    325
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Trp Asn Glu Val Asp Pro Thr Thr Asn Ser Ala Arg Val Tyr Asn
                                     340
                335
Gly Ala Ala Asp Ala Ala Ser Thr Leu Leu Gly Ala Ile Thr Ser
                350
                                     355
                                                          360
Phe Ala Ala Gly Phe Val Lys Ile Arg Trp Ala Arg Trp Ser Lys
                                                          375
                                     370
                365
Leu Leu Ile Ala Gly Val Thr Ala Thr Gln Ala Gly Leu Val Phe
                                     385
                                                          390
                380
Leu Leu Ala His Thr Arg His Pro Ser Ser Ile Trp Leu Cys Tyr
                                     400
                                                          405
                395
Ala Ala Phe Val Leu Phe Arg Gly Ser Tyr Gln Phe Leu Val Pro
                                     415
                                                          420
                410
Ile Ala Thr Phe Gln Ile Ala Ser Ser Leu Ser Lys Glu Leu Cys
                                     430
                                                          435
                425
Ala Leu Val Phe Gly Val Asn Thr Phe Phe Ala Thr Ile Val Lys
                440
                                     445
                                                          450
Thr Ile Ile Thr Phe Ile Val Ser Asp Val Arg Gly Leu Gly Leu
                                     460
                                                          465
                455
Pro Val Arg Lys Gln Phe Gln Leu Tyr Ser Val Tyr Phe Leu Ile
                                     475
                470
Leu Ser Ile Ile Tyr Phe Leu Gly Ala Met Leu Asp Gly Leu Arg
                485
                                     490
                                                          495
His Cys Gln Arg Gly His His Pro Arg Gln Pro Pro Ala Gln Gly
                                     505
                                                          510
                500
Leu Arg Ser Ala Ala Glu Glu Lys Ala Ala Gln Ala Leu Ser Val
                                     520
                                                          525
                515
Gln Asp Lys Gly Leu Gly Gly Leu Gln Pro Ala Gln Ser Pro Pro
                                                          540
                530
                                     535
Leu Ser Pro Glu Asp Ser Leu Gly Ala Val Gly Pro Ala Ser Leu
                                     550
                545
Glu Gln Arg Gln Ser Asp Pro Tyr Leu Ala Gln Ala Pro Ala Pro
                                                          570
                560
                                     565
Gln Ala Ala Glu Phe Leu Ser Pro Val Thr Thr Pro Ser Pro Cys
                575
                                     580
Thr Leu Cys Ser Ala Gln Ala Ser Gly Pro Glu Ala Ala Asp Glu
                                     595
                590
Thr Cys Pro Gln Leu Ala Val His Pro Pro Gly Val Ser Lys Leu
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Gly Leu Gln Cys Leu Pro Ser Asp Gly Val Gln Asn Val Asn Gln
                620
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<210> 168
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## <400> 168

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        Arg Met Pro
        Phe Met Trp Leu Glu Ser Gly Ile Pro Asn Leu Gly 1
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        10
        15

        Val Trp Pro Asn Arg Ile His Thr Thr Ala Glu Lys Tyr Arg Glu 20
        25
        30

        Tyr Glu Ala Arg Glu Gln Thr Asp Gln Thr Gln Ala Gln Glu Leu 35
        40
        45

        His Arg Ser Gln Asp Arg Asp Phe Glu Thr Met Ala Lys Leu His 50
        55
        60

        Ile Pro Val Met Val Asp Glu Val Val His Cys Leu Ser Pro Gln 65
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        75

        Lys Gly Gln Ile Phe Leu Asp Met Thr Phe Gly Ser Gly Gly His
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<213> Homo sapiens

<220>

<221> misc_feature

<223> Incyte ID No: LG:332027.9.orf3:2002JAN18

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                                      100
 Leu Asp Arg Asp Pro Thr Ala Tyr Ala Leu Ala Glu His Leu Ser
                 110
                                      115
                                                          120
 Glu Leu Tyr Pro Lys Gln Ile Arg Ala Met Leu Gly Gln Phe Ser
                 125
                                      130
 Gln Ala Glu Ala Leu Leu Met Lys Ala Gly Val Gln Pro Gly Thr
                 140
                                     145
 Phe Asp Gly Val Leu Met Asp Leu Gly Cys Ser Ser Met Gln Leu
                 155
                                      160
Asp Thr Pro Glu Arg Gly Phe Ser Leu Arg Lys Asp Gly Pro Leu
                 170
                                      175
Asp Met Arg Met Asp Gly Gly Arg Tyr Pro Asp Met Pro Thr Ala
                 185
                                     190
                                                          195
Ala Asp Val Val Asn Ala Leu Asp Gln Gln Ala Leu Ala Ser Ile
                 200
                                     205
Leu Arg Thr Tyr Gly Glu Glu Lys His Ala Lys Lys Ile Ala Ser
                 215
                                     220
Ala Ile Val Gln Ala Arg Ser Ile Tyr Pro Ile Thr Arg Thr Gln
                 230
                                     235
                                                          240
Gln Leu Ala Ser Ile Val Ala Gly Ala Phe Pro Pro Ser Ala Ile
                 245
                                     250
Tyr Thr Arg Lys Asp Leu Leu Gln Arg Ser Thr His Ile Ala Thr
                 260
                                     265
                                                          270
Lys Thr Phe Gln Ala Leu Arg Ile Phe Val Asn Asn Glu Leu Asn
                 275
                                     280
Glu Leu Tyr Thr Gly Leu Lys Thr Ala Gln Lys Phe Leu Arg Pro
                 290
                                     295
Gly Gly Arg Leu Val Ala Leu Ser Phe His Ser Leu Glu Asp Arg
                 305
                                     310
                                                          315
Ile Val Lys Arg Phe Leu Leu Gly Ile Ser Met Thr Glu Arg Phe
                 320
                                     325
Asn Leu Ser Val Arg Gln Gln Val Met Lys Thr Ser Gln Leu Gly
                 335
                                     340
Ser Asp His Glu Asn Thr Glu Glu Val Ser Met Arg Arg Ala Pro
                 350
                                     355
Leu Met Trp Glu Leu Ile His Lys Lys Val Leu Ser Pro Gln Asp
                 365
                                     370
Gln Asp Val Gln Asp Asn Pro Gln Arg Ala Leu Ser Gln Ala
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Ser Gly Cys Ser Leu Gln Leu Ser Pro Glu Ser Leu Lys Arg Glu
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                                      25
Pro Ala Ser Cys Leu Pro Gly Ala Met Glu Ala Val Glu Leu Ala
                 35
                                      40
Arg Lys Leu Gln Glu Glu Ala Thr Cys Ser Ile Cys Leu Asp Tyr
                 50
                                      55
Phe Thr Asp Pro Val Met Thr Thr Cys Gly His Asn Phe Cys Arg
                 65
                                      70
Ala Cys Ile Gln Leu Ser Trp Glu Lys Ala Arg Gly Lys Lys Gly
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80
                                     85
Arg Arg Lys Arg Lys Gly Ser Phe Pro Cys Pro Glu Cys Arg Glu
                                    100
                 95
Met Ser Pro Gln Arg Asn Leu Leu Pro Asn Arg Leu Leu Thr Lys
                                                         120
                110
                                    115
Val Ala Glu Met Ala Gln Gln His Pro Gly Leu Gln Lys Gln Asp
                                                         135
                                     130
                125
Leu Cys Gln Glu His His Glu Pro Leu Lys Leu Phe Cys Gln Lys
                                     145
                140
Asp Gln Ser Pro Ile Cys Val Val Cys Arg Glu Ser Arg Glu His
                                     160
                155
Arg Leu His Arg Val Leu Pro Ala Glu Glu Ala Val Gln Gly Tyr
                                                         180
                170
                                     175
Lys Leu Lys Leu Glu Glu Asp Met Glu Tyr Leu Arg Glu Gln Ile
                                                         195
                185
                                    190
Thr Arg Thr Gly Asn Leu Gln Ala Arg Glu Glu Gln Ser Leu Ala
                                     205
                                                         210
                200
Glu Trp Gln Gly Lys Val Asn Gly Ala Glu Arg Thr His Cys Ala
                                                         225
                215
                                     220
Gly Val Glu Lys Met Asn Leu Tyr Leu Val Glu Glu Glu Gln Arg
                230
                                     235
Leu Leu Gln Ala Leu Glu Thr Glu Glu Glu Glu Thr Ala Ser Arg
                                     250
                                                         255
                245
Leu Arg Glu Ser Val Ala Cys Leu Asp Arg Gln Gly His Ser Leu
                260
                                     265
Glu Leu Leu Leu Gln Leu Glu Glu Arg Ser Thr Gln Gly Pro
                                     280
                275
Leu Gln Met Leu Gln Asp Met Lys Glu Pro Leu Ser Arg Lys Asn
                                                         300
                290
                                     295
Asn Val Ser Val Gln Cys Pro Glu Val Ala Pro Pro Thr Arg Pro
                                     310
                305
Arg Thr Val Cys Arg Val Pro Gly Gln Ile Glu Val Leu Arg Gly
                                     325
                                                          330
                320
Phe Leu Gly Lys Trp Ala Pro Arg Ala Arg Thr Ser Asp Pro Gly
                                                          345
                335
                                     340
Ser Leu Gly Asp Ala Pro Leu Tyr Pro Leu Ala Ser Glu Ala Thr
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Asn Gly Gly Ser Thr Ser Ala Leu Pro Gly Asp Gly His Trp
                                                          375
                365
                                     370
Leu Phe Thr Val Pro Ser
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Ser Ser Leu Asn Thr Val Leu Ser Glu Asn Ala Arg Asp Ser Ser
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                                      25
Phe Ile Pro Leu Gly His Met Leu Thr Gln Lys Ile Ala Tyr Gln
                                      40
                  35
Ile Ala Ser Gly Leu Ala Tyr Leu His Lys Lys Asn Ile Ile Phe
                  50
                                      55
Cys Asp Leu Lys Ser Asp Asn Ile Leu Val Trp Ser Leu Asp Val
                  65
                                      70
Lys Glu His Ile Asn Ile Lys Leu Ser Asp Tyr Gly Ile Ser Arg
```

				80					85					90
Gln	Ser	Phe	His	61u 95	Gly	Ala	Leu	Gly	Val 100	Glu	Gly	Thr	Pro	Gly 105
Tyr	Glr	Ala	Pro	Glu 110	Ile	Arg	Pro	Arg	11e 115	· Val	Tyr	Asp	Glu	Lys 120
Val	Asp	Met	Phe	Ser 125	Tyr	Gly	Met	Val	Leu 130	Tyr	Glu	Leu	Leu	Ser 135
				Ala 140					Gln 145	Leu				Lys 150
				Gly 155					160					Glu 165
				Arg 170					175					Asp 180
				Lys 185					190					Gln 195
				Thr 200					205					210
				Ala 215					220					225
				Asp 230					235					240
				Lys 245 Val					250					255
				260 Glu					265					270
				275 Leu					280					285
				290 Cys					295					300
				305 Leu					310					315
				320 Arg					325					330
				335 Ala					340 Phe					345
Asp	Ala	Arg	Gln	350 Asn	Pro	Tyr	Pro	Val		Ala	Met	Glu	Val	
Asn	Ser	Gly	Ser	365 Glu 380	Val	Trp	Tyr	Ser		Gly	Pro	Gly	Leu	
Val	Ile	Asp	Cys	Ala 395	Ser	Leu	Glu	Ile	385 Cys 400	Arg	Arg	Leu	Glu	
Tyr	Met	Ala	Pro	Ser 410	Met	Val	Thr	Ser	Val 415	Val	Cys	Ser	Ser	
				Glu 425					Leu 430					435
				Tyr 440					Tyr 445					Arg
				Val 455					460					Val 465
				Thr 470					475					480
				Glu 485					490					495
				Leu 500		-			505					510
				Cys 515					520					525
				Arg 530 Val					535					540
DGI	JUL	J-01	DET	Val 545	FIO	riie	ser	mr	550	cys	GIU	Asp	ser	Asp 555

```
Met Leu His Met Pro Gly Ala Ala Ser Asp Arg Ser Glu His Asp
                560
                                     565
Leu Thr Pro Met Asp Gly Glu Thr Phe Ser Gln His Leu Gln Ala
                                     580
                575
Val Lys Ile Leu Ala Val Arg Asp Leu Ile Trp Val Pro Arg Arg
                                     595
                590
Gly Gly Asp Val Ile Val Ile Gly Leu Glu Lys Asp Ser Glu Ala
                                     610
                605
Gln Arg Gly Arg Val Ile Ala Val Leu Lys Ala Arg Glu Leu Thr
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Pro His Gly Ile Met Pro Val Ser Ser Val Lys Val Cys Trp Ala
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                635
Gly Trp Pro Val Arg Asp Met Val Tyr Met Ala Ala Val Met
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Ser Phe Gly Ile Phe Asn Leu Lys Cys Phe Pro Leu Val Val Gly
                                      25
                  20
Phe Ile Leu Pro Leu Pro Leu Pro Phe Ser Tyr Tyr Ser Glu Tyr
                                      40
Lys Pro Ala Lys Leu Ser Gln Ile Arg Gln Ile Tyr His Thr Glu
                                      55
                  50
Leu Glu Lys Tyr Glu Gln Ala Cys Asn Glu Phe Thr Thr Leu Val
                                      70
                  65
Met Asn Leu Leu Arg Glu Gln Ser Arg Thr Arg Pro Ile Ser Pro
                  80
                                      85
Lys Glu Ile Glu Arg Met Val Ser Ile Ile His Arg Lys Phe Ser
                  95
                                     100
Ser Ile Gln Met Gln Leu Lys Gln Ser Thr Cys Glu Ala Val Met
                 110
                                     115
                                                          120
Ile Leu Arg Ser Arg Phe Leu Asp Ala Arg Arg Lys Arg Asn
                                     130
                 125
 Phe Asn Lys Gln Ala Thr Glu Ile Leu Asn Glu Tyr Phe Tyr Ser
                 140
                                     145
His Leu Ser Asn Pro Tyr Pro Ser Glu Glu Ala Lys Glu Glu Leu
                 155
                                     160
 Ala Lys Lys Cys Gly Ile Thr Val Ser Gln Val Ser Asn Trp Phe
                                      175
                 170
 Gly Asn Lys Arg Ile Arg Tyr Lys Lys Asn Ile Gly Lys Phe Gln
                 185
                                      190
 Glu Glu Ala Asn Ile Tyr Cys Cys Gln Asn Ser Cys His Cys Tyr
                 200
                                      205
 Gln Cys Val Ser Pro Trp Lys Pro Ser
                 215
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 1
Thr Cys Ala Ile Cys Leu Gly Leu Tyr Gln Asp Pro Val Thr Leu
                 20
Pro Cys Gly His Asn Phe Cys Gly Ala Cys Ile Arg Asp Trp Trp
                 35
                                      40
Asp Arg Cys Gly Lys Ala Cys Pro Glu Cys Arg Glu Pro Phe Pro
                                      55
                 50
Asp Gly Ala Glu Leu Arg Arg Asn Val Ala Leu Ser Gly Val Leu
                                      70
                 65
Glu Val Val Arg Ala Gly Pro Ala Arg Asp Pro Gly Pro Asp Pro
                 80
                                      85
Gly Pro Gly Pro Asp Pro Ala Ala Arg Cys Pro Arg His Gly Arg
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                 95
Pro
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Ser Ser Pro Ser Cys Cys Arg Tyr Arg Arg Cys Cys Arg Arg Leu
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Arg Pro Pro Leu Arg Ser Val Val Gln Pro Gly Pro Arg Thr Met
                 35
                                      40
Ser Leu Ser Arg Ser Glu Glu Met His Arg Leu Thr Glu Asn Val
                 50
                                      55
Tyr Lys Thr Ile Met Glu Gln Phe Asn Pro Ser Leu Arg Asn Phe
                                      70
                  65
Ile Ala Met Gly Lys Asn Tyr Glu Lys Ala Leu Ala Gly Val Thr
                                      85
                 80
Tyr Ala Ala Lys Gly Tyr Phe Asp Ala Leu Val Lys Met Gly Glu
                 95
                                     100
Leu Ala Ser Glu Ser Gln Gly Ser Lys Glu Leu Gly Asp Val Leu
                 110
                                     115
Phe Gln Met Ala Glu Val His Arg Gln Ile Gln Asn Gln Leu Glu
                                     130
                125
Glu Met Leu Lys Ser Phe His Asn Glu Leu Leu Thr Gln Leu Glu
                                                          150
                 140
                                     145
Gln Lys Val Glu Leu Asp Ser Arg Tyr Leu Ser Ala Ala Leu Lys
                                                          165
                 155
                                     160
Lys Tyr Gln Thr Glu Gln Arg Ser Lys Gly Asp Ala Leu Asp Lys
                                                          180
                 170
                                     175
Cys Gln Ala Glu Leu Lys Lys Leu Arg Lys Lys Ser Gln Gly Ser
                 185
                                     190
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Lys Asn Pro Gln Lys Tyr Ser Asp Lys Glu Leu Gln Tyr Ile Asp
                 200
                                      205
Ala Ile Ser Asn Lys Gln Gly Glu Leu Glu Asn Tyr Val Ser Asp
                 215
                                     220
Gly Tyr Lys Thr Ala Leu Thr Glu Glu Arg Arg Arg Phe Cys Phe
                 230
                                     235
Leu Val Glu Lys Gln Cys Ala Val Ala Lys Asn Ser Ala Ala Tyr
                 245
                                     250
His Ser Lys Gly Lys Glu Leu Leu Ala Gln Lys Leu Pro Leu Trp
                 260
                                     265
                                                          270
Gln Gln Ala Cys Ala Asp Pro Ser Lys Ile Pro Glu Arg Ala Val
                 275
                                     280
Gln Leu Met Gln Gln Val Ala Ser Asn Gly Ala Thr Leu Pro Ser
                 290
                                     295
Ala Cys Arg Pro Pro Ser Gln Pro Gly His Phe Arg Pro His Ser
                 305
                                     310
                                                          315
Gly Gly Gln Ala Pro Ala Gly Ala Pro Arg Ala Gly Thr Val Arg
                 320
                                     325
Gly Ala Asp Val Cys Pro Gly Glu His Thr His His Glu Arg Arg
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                                     340
His Arg Pro Gly Trp Arg Gly Leu Gln Pro Val Gly
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Met Ala Leu Phe Ser Cys Arg Asn Ala Val Glu Glu Gly Lys Gly
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Ile Phe Tyr Asn Ile Lys Asn Phe Val Arg Phe Gln Leu Ser Thr
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Ser Ile Ser Ala Leu Ser Leu Ile Thr Leu Ser Thr Val Phe Asn
                 35
                                      40
Leu Pro Ser Pro Leu Asn Ala Met Gln Ile Leu Trp Ile Asn Ile
                 50
                                      55
Ile Met Asp Gly Pro Pro Ala Gln Arg Ser Ser Gln Lys Thr Glu
                 65
                                      70
Val Cys Cys Thr Gly Val Arg Leu Gly Val Glu Gly Arg Gly Glu
                 80
                                      85
Ser Thr Trp Ala Gly Arg Ala Gly Leu Gly Val Glu Pro Val Asp
                 95
                                     100
Lys Asp Ala Phe Arg Gln Pro Pro Arg Ser Val Arg Asp Thr Ile
                110
                                     115
                                                         120
Leu Ser Arg Ala Leu Ile Leu Lys Ile Leu Met Ser Ala Ala Ile
                125
                                     130
Ile Ile Ser Gly Thr Leu Phe Ile Phe Trp Lys Glu Met Pro Glu
                140
                                     145
                                                         150
Asp Arg Ala Ser Thr Pro Arg Thr Thr Thr Met Thr Phe Thr Cys
                155
                                     160
Phe Val Phe Phe Asp Leu Phe Asn Ala Leu Thr Cys Arg Ser Gln
                170
                                     175
Thr Lys Leu Ile Phe Glu Ile Gly Phe Leu Arg Asn His Met Phe
                185
                                     190
                                                         195
Leu Tyr Ser Val Leu Gly Ser Ile Leu Gly Gln Leu Ala Val Ile
                200
                                     205
                                                         210
Tyr Ile Pro Pro Leu Gln Arg Val Phe Gln Thr Glu Asn Leu Gly
                215
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Ala Leu Asp Leu Leu Phe Leu Thr Gly Leu Ala Ser Ser Val Phe
                230
                                     235
Ile Leu Ser Glu Leu Leu Lys Leu Cys Glu Lys Tyr Cys Cys Ser
                                     250
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Pro Lys Arg Val Gln Met His Pro Glu Asp Val
                260
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Pro Pro Gly Val Pro Val Ser Asn Val Asn Leu Glu Ile Arg Pro
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                                      25
Thr Gly Gly Gln Leu Ile Glu Gly Glu Asn Met Val Leu Ile Cys
                                                           45
                                      40
                 35
Ser Val Ala Gln Gly Ser Gly Thr Val Thr Phe Ser Trp His Lys
                                                           60
                 50
                                      55
Glu Gly Arg Val Arg Ser Leu Gly Arg Lys Thr Gln Arg Ser Leu
                                      70
                 65
Leu Ala Glu Leu His Val Leu Thr Val Lys Glu Ser Asp Ala Gly
                                      85
                 80
Arg Tyr Tyr Cys Ala Ala Asp Asn Val His Ser Pro Ile Leu Ser
                                     100
Thr Trp Ile Arg Val Thr Val Arg Ile Pro Val Ser His Pro Val
                                     115
                110
Leu Thr Phe Arg Ala Pro Arg Ala His Thr Val Val Gly Asp Leu
                                                          135
                125
                                     130
Leu Glu Leu His Cys Glu Ser Leu Arg Gly Ser Pro Pro Ile Leu
                                     145
                140
Tyr Arg Phe Tyr His Glu Asp Val Thr Leu Gly Asn Ser Ser Ala
                                     160
                155
Pro Ser Gly Gly Ala Ser Phe Asn Leu Ser Leu Thr Ala Glu
                 170
                                     175
His Ser Gly Asn Tyr Ser Cys Asp Ala Asp Asn Gly Leu Gly Ala
                                     190
                 185
Gln His Ser His Gly Val Ser Leu Arg Val Thr Val Pro Val Ser
                 200
                                     205
Arg Pro Val Leu Thr Leu Arg Ala Pro Gly Ala Gln Ala Val Val
                 215
                                     220
Gly Asp Leu Leu Glu Leu His Cys Glu Ser Leu Arg Gly Ser Phe
                                     235
                                                          240
                 230
Pro Ile Leu Tyr Trp Phe Tyr His Glu Asp Asp Thr Leu Gly Asn
                                     250
                 245
Ile Ser Ala His Ser Gly Gly Gly Ala Ser Phe Asn Leu Ser Leu
                                                          270
                                     265
                 260
Thr Thr Glu His Ser Gly Asn Tyr Ser Cys Glu Ala Asp Asn Gly
                 275
                                     280
                                                          285
Leu Gly Ala Gln His Ser Lys Val Val Thr Leu Asn Val Thr Gly
                                     295
                                                          300
                 290
Thr Ser Arg Asn Arg Thr Gly Leu Thr Ala Ala Gly Ile Thr Gly
                                      310
                                                          315
                 305
Leu Val Leu Ser Ile Leu Val Leu Ala Ala Ala Ala Leu Leu
                                                          330
                 320
                                      325
His Tyr Ala Arg Ala Arg Arg Lys Pro Gly Gly Leu Ser Ala Thr
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                 335
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Gly Thr Ser Ser His Ser Pro Ser Glu Cys Gln Glu Pro Ser Ser
                350
                                     355
Ser Arg Pro Ser Arg Ile Asp Pro Gln Glu Pro Thr His Ser Lys
                                                         375
                365
                                     370
Pro Leu Ala Pro Met Glu Leu Glu Pro Met Tyr Ser Asn Val Asn
                380
                                     385
Pro Gly Asp Ser Asn Pro Ile Tyr Ser Gln Ile Trp Ser Ile Gln
                395
                                     400
His Thr Lys Glu Asn Ser Ala Asn Cys Pro Met Met His Gln Glu
                410
                                     415
                                                         420
His Glu Glu Leu Thr Val Leu Tyr Ser Glu Leu Lys Lys Thr His
                425
                                     430
Pro Asp Asp Ser Ala Gly Glu Ala Ser Ser Arg Gly Arg Ala His
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Glu Glu Asp Asp Glu Glu Asn Tyr Glu Asn Val Pro Arg Val Leu
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Leu Ala Ser Asp His
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His Arg Gly Ser Pro Ile Lys Leu Val Asn Ile Asn Ser Thr Asp
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                 20
Ile Ala Asp Gly Arg Pro Ser Ile Val Leu Gly Leu Met Trp Thr
                  35
                                      40
Ile Ile Leu Tyr Phe Gln Ile Glu Glu Leu Thr Ser Asn Leu Pro
                  50
                                      55
Gln Leu Gln Ser Leu Ser Ser Ser Ala Ser Ser Val Asp Ser Ile
                  65
                                      70
Val Ser Ser Glu Thr Pro Ser Pro Pro Ser Lys Arg Lys Val Thr
                  80
                                      85
Thr Lys Ile Gln Gly Asn Ala Lys Lys Ala Leu Leu Lys Trp Val
                  95
                                     100
                                                          105
Gln Tyr Thr Ala Gly Lys Gln Thr Gly Ile Glu Val Lys Asp Phe
                 110
                                     115
                                                          120
Gly Lys Ser Trp Arg Ser Gly Val Ala Phe His Ser Val Ile His
                 125
                                     130
Ala Ile Arg Pro Glu Leu Val Asp Leu Glu Thr Val Lys Gly Arg
                 140
                                                          150
                                     145
Ser Asn Arg Glu Asn Leu Glu Asp Ala Phe Thr Ile Ala Glu Thr
                 155
                                     160
Glu Leu Gly Ile Pro Arg Leu Leu Asp Pro Glu Asp Val Asp Val
                 170
                                     175
Asp Lys Pro Asp Glu Lys Ser Ile Met Thr Tyr Val Ala Gln Phe
                                     190
                 185
Leu Lys His Tyr Pro Asp Ile His Asn Ala Ser Thr Asp Gly Gln
                 200
                                     205
Glu Asp Asp Glu Ile Leu Pro Gly Phe Pro Ser Phe Ala Asn Ser
                 215
                                     220
Val Gln Asn Phe Lys Arg Glu Asp Arg Val Ile Phe Lys Glu Met
                 230
                                     235
Lys Val Trp Ile Glu Gln Phe Glu Arg Asp Leu Thr Arg Ala Gln
                 245
                                     250
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Met	Val	Glu	Ser	Asn 260	Leu	Gln	Asp	Lys	Tyr 265	Gln	Ser	Phe	Lys	His 270
Phe	Arg	Val	Gln		Glu	Met	Lys	Arg		Gln	Ile	Glu	His	
Ile	Gln	Pro	Leu		Arg	Asp	Gly	Lys		Ser	Leu	Asp	Gln	
Leu	Val	Lys	Gln		Trp	Asp	Arg	Va1		Ser	Arg	Leu	Phe	
Trp	His	Ile	Gl'n		Asp	Lys	Ser	Leu		Ala	Pro	Leu	Gly	
Ile	Gly	Ala	Trp	Leu	Tyr	Arg	Ala	Glu		Ala	Leu	Arg	Glu	
Ile	Thr	Val	Gln	335 Gln 350	Val	His	Glu	Glu		Ala	Asn	Thr	Ile	
Arg	Lys	Leu	Glu		His	Lys	Asp	Leu		Gln	Asn	Thr	Asp	
His	Lys	Arg	Ala		His	Glu	Ile	Tyr		Thr	Arg	Ser	Val	
Gly	Ile	Pro	Val		Pro	Asp	Gln	Leu		Asp	Met	Ala	Glu	
Phe	His	Phe	Val		Ser	Thr	Ser	Glu		His	Leu	Met	Lys	
Glu	Phe	Leu	Glu		Lys	Tyr	Arg	Leu		Ser	Leu	Leu	Val	
Ala	Glu	Ser	Lys	Leu	Lys	Ser	Trp	Ile		Lys	Tyr	Gly	Arg	
Glu	Ser	Val	Glu	440 Gln 455	Leu	Leu	Gln	Asn		Val	Ser	Phe	Ile	
Asn	Ser	Lys	Phe	Phe		Gln	Tyr	Glu		Thr	Tyr	Gln	Ile	
Lys	Gln	Thr	Ala		Met	Tyr	Val	Lys		Asp	Gly	Ser	Val	
Glu	Ala	Glu	Asn		Met	Lys	Phe	Met	Asn	Glu	Thr	Thr	Ala	
Trp	Arg	Asn	Leu		Val	Glu	Val	Arg		Val	Arg	Ser	Met	
Glu	Glu	Val	Ile		Asn	Trp	Asp	Arg		Gly	Asn	Thr	Val	Ala
Ser	Leu	Gln	Ala		Leu	Glu	Asp	Ala			Met	Leu	Asn	540 Gln
Ser	Glu	Asn	Ala		Lys	Asp	Phe	Phe		Asn	Leu	Pro	His	555 Trp
Ile	Gln	Gln	His		Ala	Met	. Asn	Asp		Gly	Asn	Phe	Leu	
Glu	Thr	Cys	Asp		Met	Val	Ser	Arg		Leu	Lys	Gln	Gln	585 Leu
Leu	Leu	Leu	Asn	590 Gly		Trp	Arg	Glu		Phe	Met	Glu	Val	600 Lys
Gln	Tyr	Ala	Gln	605 Ala		Glu	. Met	Asp		Met	Lys	Lys	Glu	615 Tyr
Thr	Asp	Сув	. Val	620 Val		Lev	Ser	Ala		Ala	Thr	Glu	Ala	630 His
Lys	Lys	Leu	Ser	635 Glu		Leu	ı Glu	Val		Phe	Met	. Asn	Val	645 Lys
Leu	Leu	ıle	Gln	650 Asp		Glu	ı Asp	Ile	655 Glu		Arg	r Val	Pro	660 Val
Met	Asp	Ala	ı Glm	665 Tyr		: Ile	e Ile	Thr		Thr	Ala	His	Leu	675
Thr	Lys	Glu	ser	680 Pro		Glu	ı Glu	Gly		Glu	Met	: Phe	a Ala	690 Thr
				695	5				700	)				705 Tyr
				710	)				715	5				720 Glu
				_										

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725
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Glu Leu Glu Lys Gln Met Thr Ser Phe Tyr Asp Ser Leu Gly Lys
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Ile Asn Glu Ile Ile Thr Val Leu Glu Arg Glu Ala Gln Ser Ser
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                                     760
Ala Leu Phe Lys Gln Lys His Gln Glu Leu Leu Ala Cys Gln Glu
                770
                                     775
Asn Cys Lys Lys Thr Leu Thr Leu Ile Glu Lys Gly Ser Gln Ser
                785
                                     790
Val Gln Lys Phe Val Thr Leu Ser Asn Val Leu Lys His Phe Asp
                800
                                     805
Gln Thr Arg Leu Gln Arg Gln Ile Ala Asp Ile His Val Ala Phe
                815
                                     820
Gln Ser Met Val Lys Lys Thr Gly Asp Trp Lys Lys His Val Glu
                830
                                     835
Thr Asn Ser Arg Leu Met Lys Lys Phe Glu Glu Ser Arg Ala Glu
                845
                                     850
Leu Glu Lys Val Leu Arg Ile Ala Gln Glu Gly Leu Glu Glu Lys
                860
                                     865
                                                         870
Gly Asp Pro Glu Glu Leu Leu Arg Arg His Thr Glu Phe Phe Ser
                875
                                     880
Gln Leu Asp Gln Arg Val Leu Asn Ala Phe Leu Lys Ala Cys Asp
                890
                                     895
Glu Leu Thr Asp Ile Phe Gln Ser Arg Ser Ser Arg Gly Cys Arg
                905
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Lys Leu Phe Glu Ser Ser Thr Asn Asn Gly Arg Ile Phe Lys Glu
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Lys Pro Leu Ile Ile Cys Phe Ile
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His Leu Lys Thr Met Val Ile Glu Asn Leu Glu Gly Asn Lys His
                 35
                                     40
Ile Thr His Val Asp Leu Arg Asp Asn Arg Leu Thr Asp Leu Asp
                 50
                                     55
Leu Ser Ser Leu Cys Ser Leu Glu Gln Leu His Cys Gly Arg Asn
                 65
                                      70
Gln Leu Arg Glu Leu Thr Leu Ser Gly Phe Ser Leu Arg Thr Leu
                                      85
Tyr Ala Ser Ser Asn Arg Leu Thr Ala Val Asn Val Tyr Pro Val
                 95
                                     100
Pro Ser Leu Leu Thr Phe Leu Asp Leu Ser Arg Asn Leu Leu Glu
                110
                                    115
Cys Val Pro Asp Trp Ala Cys Glu Ala Lys Lys Ile Glu Val Leu
                125
                                    130
Asp Val Ser Tyr Asn Leu Leu Thr Glu Val Pro Val Arg Ile Leu
                140
                                    145
Ser Ser Leu Ser Leu Arg Lys Leu Met Leu Gly His Asn His Val
                155
                                    160
Gln Asn Leu Pro Thr Leu Val Glu His Ile Pro Leu Glu Val Leu
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				170					175					180
				185			Thr		190					195
Ser	Lys	Ala	Leu	Asn 200	Leu	Arg	Tyr	Leu	Asn 205	Ala	Ser	Ala	Asn	Ser 210
Leu	Glu	Ser	Leu		Ser	Ala	Cys	Thr	Gly 220	Glu	Glu	Ser	Leu	Ser 225
Met	Leu	Gln	Leu		Tyr	Leu	Thr	Asn		Leu	Leu	Thr	Asp	
Cys	Ile	Pro	Val		Val	Gly	His	Leu		Leu	Arg	Ile	Leu	
Leu	Ala	Asn	Asn		Leu	Gln	Thr	Phe		Ala	Ser	Lys	Leu	
Lys	Leu	Glu	Gln	Leu 275	Glu	Glu	Leu	Asn		Ser	Gly	Asn	Lys	
Lys	Thr	Ile	Pro		Thr	Ile	Ala	Asn		Lys	Arg	Leu	His	
Leu	Val	Ala	His		Àsn	Asn	Ile	Ser		Phe	Pro	Glu	Ile	
Gln	Leu	Pro	Gln	Ile 320	Gln	Phe	Val	Asp		Ser	Cys	Asn	Asp	
Thr	Glu	Ile	Leu		Pro	Glu	Ala	Leu		Ala	Thr	Leu	Gln	
Leu	Asp	Leu	Thr		Asn	Thr	Asn	Leu		Leu	Glu	His	Lys	Thr 360
Leu	Asp	Ile	Phe		His	Ile	Thr	Thr		Lys	Ile	Asp	Gln	
Pro	Leu	Pro	Thr		Asp	Ser	Thr	Val		Ser	Thr	Phe	Trp	
His	Gly	Leu	Ala		Met	Ala	Gly	Gln		Asn	Lys	Leu	Cys	
Ser	Ala	Leu	Ala		Asp	Ser	Phe	Ala		Gly	Val	Gly	Ala	
Туг	Gly	Met	Phe			Asp	Arg	Asn		Glu	Leu	Pro	Arg	
Leu	Gln	Cys	Thr		Ala	Asp	Val	Leu		Glu	Glu	Val	Gln	
Ser	Thr	Asn	Asp		Val	Phe	Met	Ala			Phe	Leu	Val	
His	Arg	Lys	Leu		Met	Ala	Gly	Gln		Leu	Gly	Ser	Ser	
Leu	Leu	Суя	Tyr		Arg	Pro	Asp	Thr		Asp	Pro	Ala	Ser	
Phe	Ser	Lev	Thr		Ala	Asn	val	Gly		Суѕ	Gln	Ala	. Val	
Сув	Arg	Gly	gly		Pro	Val	Pro	Lev		Lys	Val	Phe	e Ser	Leu 525
Glu	Gln	Asp	Pro		Glu	Ala	a Gln	Arg		Lys	Asp	Glr	Lys	Ala 540
Ile	Ile	Thr	Glu		Asr	Lys	Val	Asr		Val	Thr	Сув	cys	Thr 555
Arg	Met	. Let	ı Gly		Thr	Tyr	Leu	ι Туг		Trp	ıle	e Lev	Pro	Lys 570
Pro	His	: Ile	e Ser		Thr	Pro	Lev	Thr		Glr	Asp	Gli	ı Lev	Leu 585
Ile	Lev	ı Gly	/ Asr		Ala	Let	ı Trp	Glu		Leu	ser	туз	Thr	Glu 600
Ala	. Val	Ası	n Ala		Arg	y His	s Val	Glr		Pro	Let	ı Ala	a Ala	Ala 615
Lys	Lys	Le	л Суя		Le	ı Ala	a Glr	ı Sei		Gl3	у Суз	s Glr	a Asp	Asn 630
Val	. G13	, Ala	a Met		L Vai	l Ty	r Lei	ı Ası		• Gl3	/ Glu	a Glu	ı Gly	Cys 645
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Thr Cys Glu Met Asn Gly Leu Thr Leu Pro Gly Pro Val Gly Phe
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 Ala Ser Thr Thr Thr Ile Lys Asp Ala Pro Lys Pro Ala Thr Pro
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                                      670
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 Ser Ser Ser Gly Ile Ala Ser Glu Phe Ser Ser Glu Met Ser
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                                      685
 Thr Ser Glu Val Ser Ser Glu Val Gly Ser Thr Ala Ser Asp Glu
                                      700
                                                           705
 His Asn Ala Gly Gly Leu Asp Thr Ala Leu Leu Pro Arg Pro Glu
                  710
                                      715
 Arg Arg Cys Ser Leu His Pro Thr Pro Thr Ser Gly Leu Phe Gln
                  725
                                      730
                                                           735
 Arg Gln Pro Ser Ser Ala Thr Phe Ser Ser Asn Gln Ser Asp Asn
                  740
                                      745
 Gly Leu Asp Ser Asp Asp Gln Pro Val Glu Gly Val Ile Thr
                 755
                                      760
 Asn Gly Ser Lys Val Glu Val Glu Val Asp Ile His Cys Cys Arg
                  770
                                      775
 Gly Arg Asp Leu Glu Asn Ser Pro Pro Leu Ile Glu Ser Ser Pro
                 785
                                      790
 Thr Leu Cys Ser Glu Glu His Ala Arg Gly Ser Cys Phe Gly Ile
                 800
                                      805
 Arg Arg Gln Asn Ser Val Asn Ser Gly Met Leu Leu Pro Met Ser
                 815
                                      820
 Lys Asp Arg Met Glu Leu Gln Lys Ser Pro Ser Thr Ser Cys Leu
                                      835
                                                          840
 Tyr Gly Lys Lys Leu Ser Asn Gly Ser Ile Val Pro Leu Glu Asp
                 845
                                      850
 Ser Leu Asn Leu Ile Glu Val Ala Thr Glu Val Pro Lys Arg Lys
                 860
                                      865
                                                          870
 Thr Gly Tyr Phe Ala Ala Pro Thr Gln Met Glu Pro Glu Asp Gln
                 875
                                      880
 Phe Val Val Pro His Asp Leu Glu Glu Glu Val Lys Glu Gln Met
                 890
                                     895
                                                          900
Lys Gln His Gln Asp Ser Arg Leu Glu Pro Glu Pro His Glu Glu
                 905
                                     910
                                                          915
Asp Arg Thr Glu Pro Pro Glu Glu Phe Asp Thr Ala Leu
                 920
                                     925
<210> 179
<211> 304
<212> PRT
<213> Homo sapiens
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Pro Gly Pro Trp Ala Pro Ala Arg Ala Gly Ala Gly Ala Ser Gly
                                      10
                                                          15
Met Ala Phe Arg Gln Ala Leu Gln Leu Ala Ala Cys Gly Leu Ala
                  20
                                      25
Gly Gly Ser Ala Ala Val Leu Phe Ser Ala Val Ala Val Gly Lys
                  35
                                      40
Pro Arg Ala Gly Gly Asp Ala Glu Pro Arg Pro Ala Glu Pro Pro
                  50
                                      55
Ala Trp Ala Gly Gly Ala Arg Pro Gly Pro Gly Val Trp Asp Pro
                 65
                                      70
                                                          75
Asn Trp Asp Arg Arg Glu Pro Leu Ser Leu Ile Asn Val Arg Lys
                                      85
Arg Asn Val Glu Ser Gly Glu Glu Leu Ala Ser Lys Leu Asp
```

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His Tyr Lys Ala Lys Ala Thr Arg His Ile Phe Leu Ile Arg His
                110
                                     115
Ser Gln Tyr His Val Asp Gly Ser Leu Glu Lys Asp Arg Thr Leu
                                                          135
                                     130
                125
Thr Pro Leu Gly Arg Glu Gln Ala Glu Leu Thr Gly Leu Arg Leu
                                     145
                140
Ala Ser Leu Gly Leu Lys Phe Asn Lys Ile Val His Ser Ser Met
                                     160
                155
Thr Arg Ala Ile Glu Thr Thr Asp Ile Ile Ser Arg His Leu Pro
                                     175
                170
Gly Val Cys Lys Val Ser Thr Asp Leu Leu Arg Glu Gly Ala Pro
                                     190
                185
Ile Glu Pro Asp Pro Pro Val Ser His Trp Lys Pro Glu Ala Val
                                     205
                200
Gln Tyr Tyr Glu Asp Gly Ala Arg Ile Glu Ala Ala Phe Arg Asn
                215
                                     220
Tyr Ile His Arg Ala Asp Ala Arg Gln Glu Glu Asp Ser Tyr Glu
                                     235
                230
Ile Phe Ile Cys His Ala Asn Val Ile Arg Tyr Ile Val Cys Arg
                                     250
                                                          255
                245
Ala Leu Gln Phe Pro Pro Glu Gly Trp Leu Arg Leu Ser Leu Asn
                                                          270
                260
                                     265
Asn Gly Ser Ile Thr His Leu Val Ile Arg Pro Asn Gly Arg Val
                                                          285
                                     280
                275
Ala Leu Arg Thr Leu Gly Asp Thr Gly Phe Met Pro Pro Asp Lys
                                                          300
                                     295
                290
Ile Thr Arg Ser
<210> 180
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<211> 320

<212> PRT

<213> Homo sapiens

<220>

<221> misc_feature

<223> Incyte ID No: LG:410461.92.orf3:2002JAN18

<400> 180

Ala Pro Ala Pro Glu Pro Gly Pro Arg Ala Ala Ala Ala Gly Gly Thr Met Ser Cys Ala Gly Arg Ala Gly Pro Ala Arg Leu Ala Ala Leu Ala Leu Leu Thr Cys Ser Leu Trp Pro Ala Arg Ala Asp Asn Ala Ser Gln Glu Tyr Tyr Thr Ala Leu Ile Asn Val Thr Val Gln Glu Pro Gly Arg Gly Ala Pro Leu Thr Phe Arg Ile Asp Arg Gly Arg Tyr Gly Leu Asp Ser Pro Lys Ala Glu Val Arg Gly Gln Val Leu Ala Pro Leu Pro Leu His Gly Val Ala Asp His Leu Gly Cys Asp Pro Gln Thr Arg Phe Phe Val Pro Pro Asn Ile Lys Gln Trp Ile Ala Leu Leu Gln Arg Gly Asn Cys Thr Phe Lys Glu Lys Ile Ser Arg Ala Ala Phe His Asn Ala Val Ala Val Val Ile Tyr Asn Asn Lys Ser Lys Glu Glu Pro Val Thr Met Thr His Pro Gly Thr Gly Asp Ile Ile Ala Val Met Ile Thr Glu Leu Arg Gly Lys 

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Asp Ile Leu Ser Tyr Leu Glu Asn Phe Ser Arg Gly Ser Leu Val
                 185
                                      190
 Phe Val Ser Ile Ser Phe Ile Val Leu Met Ile Ile Ser Ser Ala
                 200
                                      205
 Trp Leu Ile Phe Tyr Phe Ile Gln Lys Ile Arg Tyr Thr Asn Ala
                 215
                                      220
Arg Asp Arg Asn Gln Arg Arg Leu Gly Asp Ala Ala Lys Lys Ala
                 230
                                      235
Ile Ser Lys Leu Thr Thr Arg Thr Val Lys Lys Gly Asp Lys Glu
                 245
                                      250
Thr Asp Pro Asp Phe Asp His Cys Ala Val Cys Ile Glu Ser Tyr
                 260
                                      265
                                                          270
Lys Gln Asn Asp Val Val Arg Ile Leu Pro Cys Lys His Val Phe
                 275
                                      280
                                                          285
His Lys Ser Cys Val Asp Pro Trp Leu Ser Glu His Cys Thr Cys
                 290
                                      295
                                                          300
Pro Met Cys Lys Leu Asn Ile Leu Lys Ala Leu Gly Asn Cys Ala
                 305
                                     310
Glu Phe Ala Met Tyr
                 320
<210> 181
<211> 358
<212> PRT
<213> Homo sapiens
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<400> 181
Asn His Cys Pro Thr Arg Ala Met Ile Val Gln Arg Val Val Leu
Asn Ser Arg Pro Gly Lys Asn Gly Asn Pro Val Ala Glu Asn Phe
                  20
                                      25
Arg Met Glu Glu Val Tyr Leu Pro Asp Asn Ile Asn Glu Gly Gln
                  35
                                      40
Val Gln Val Arg Thr Leu Tyr Leu Ser Val Asp Pro Tyr Met Arg
                  50
                                      55
Cys Arg Met Asn Glu Asp Thr Gly Thr Asp Tyr Ile Thr Pro Trp
                  65
                                      70
Gln Leu Ser Gln Val Val Asp Gly Gly Gly Ile Gly Ile Ile Glu
                  80
                                      85
Glu Ser Lys His Thr Asn Leu Thr Lys Gly Asp Phe Val Thr Ser
                 95
                                     100
Phe Tyr Trp Pro Trp Gln Thr Lys Val Ile Leu Asp Gly Asn Ser
                 110
                                     115
Leu Glu Lys Val Asp Pro Gln Leu Val Asp Gly His Leu Ser Tyr
                125
                                     130
                                                          135
Phe Leu Gly Ala Ile Gly Met Pro Gly Leu Thr Ser Leu Ile Gly
                140
                                     145
                                                          150
Ile Gln Glu Lys Gly His Ile Thr Ala Gly Ser Asn Lys Thr Met
                155
                                     160
Val Val Ser Gly Ala Ala Gly Ala Cys Gly Ser Val Ala Gly Gln
                170
                                     175
                                                          180
Ile Gly His Phe Leu Gly Cys Ser Arg Val Val Gly Ile Cys Gly
                185
                                     190
Thr His Glu Lys Cys Ile Leu Leu Thr Ser Glu Leu Gly Phe Asp
                200
                                     205
Ala Ala Ile Asn Tyr Lys Lys Asp Asn Val Ala Glu Gln Leu Arg
                215
                                     220
                                                         225
Glu Ser Cys Pro Ala Gly Val Asp Val Tyr Phe Asp Asn Val Gly
                230
                                     235
                                                         240
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Gly Asn Ile Ser Asp Thr Val Ile Ser Gln Met Asn Glu Asn Ser
                                     250
His Ile Ile Leu Cys Gly Gln Ile Ser Gln Tyr Asn Lys Asp Val
                                                         270
                                     265
                260
Pro Tyr Pro Pro Pro Leu Ser Pro Ala Ile Glu Ala Ile Gln Lys
                                                          285
                                     280
                275
Glu Arg Asn Ile Thr Arg Glu Arg Phe Leu Val Leu Asn Tyr Lys
                                     295
                                                         300
Asp Lys Phe Glu Pro Gly Ile Leu Gln Leu Ser Gln Trp Phe Lys
                                                         315
                305
                                     310
Glu Gly Lys Leu Lys Ile Lys Glu Thr Val Ile Asn Gly Leu Glu
                                     325
                                                          330
                320
Asn Met Gly Ala Ala Phe Gln Ser Met Met Thr Gly Gly Asn Ile
                335
                                     340
Gly Lys Gln Ile Val Cys Ile Ser Glu Glu Ile Ser Leu
                350
<210> 182
<211> 438
<212> PRT
<213> Homo sapiens
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<400> 182
Arg Val Arg Ala Gly Leu Pro Cys Pro Met Cys Ser Gly Arg Phe
Gln Asn Ile Gln Val Asn Pro Asp Phe Pro Arg Gly Arg Ile Ser
                                      25
Asn Ser Phe Arg Arg Thr Ser Ser Thr Glu Asn Lys Thr Lys Thr
                 35
                                      40
Leu Gly Lys Leu His Gln Glu Pro Arg Gln Leu Gln Ser Asp Gly
                                      55
                  50
Lys Arg Lys Ile Leu Leu Glu Glu Leu Ala Asn Ser Asp Pro Lys
                                      70
                  65
Leu Ala Leu Thr Gly Val Pro Ile Val Gln Trp Pro Lys Arg Asp
                                      85
                                                           90
                 80
Lys Leu Lys Phe Pro Thr Arg Pro Lys Val Arg Val Pro Thr Ile
                  95
                                     100
Pro Ile Thr Lys Pro His Thr Met Lys Pro Ala Pro Arg Leu Thr
                                     115
                 110
Pro Val Arg Pro Ala Ala Ala Ser Pro Ile Val Ser Gly Ala Arg
                                      130
                                                          135
                 125
Arg Arg Arg Val Arg Cys Arg Lys Cys Lys Ala Cys Val Gln Gly
                                                          150
                                      145
                 140
Glu Cys Gly Val Cys His Tyr Cys Arg Asp Met Lys Lys Phe Gly
                 155
                                      160
Gly Pro Gly Arg Met Lys Gln Ser Cys Val Leu Arg Gln Cys Leu
                                      175
                 170
Ala Pro Arg Leu Pro His Ser Val Thr Cys Ser Leu Cys Gly Glu
                                      190
                 185
Val Asp Gln Asn Glu Glu Thr Gln Asp Phe Glu Lys Lys Leu Met
                 200
                                      205
                                                          210
Glu Cys Cys Ile Cys Asn Glu Ile Val His Pro Gly Cys Leu Gln
                                                          225
                                      220
                 215
Met Asp Gly Glu Gly Leu Leu Asn Glu Glu Leu Pro Asn Cys Trp
                                                          240
                 230
                                      235
Glu Cys Pro Lys Cys Tyr Gln Glu Asp Ser Ser Glu Lys Ala Gln
                                                          255
                 245
                                      250
Lys Arg Lys Met Glu Glu Ser Asp Glu Glu Ala Val Gln Ala Lys
                                      265
                 260
```

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Val Leu Arg Pro Leu Arg Ser Cys Asp Glu Pro Leu Thr Pro Pro
                275
                                     280
Pro His Ser Pro Thr Ser Met Leu Gln Leu Ile His Asp Pro Val
                290
                                     295
                                                         300
Ser Pro Arg Gly Met Val Thr Arg Ser Ser Pro Gly Ala Gly Pro
                305
                                     310
Ser Asp His His Ser Ala Ser Arg Asp Glu Arg Phe Lys Arg Arg
                320
                                     325
                                                         330
Gln Leu Leu Arg Leu Gln Ala Thr Glu Arg Thr Met Val Arg Glu
                335
Lys Glu Asn Asn Pro Ser Gly Lys Lys Glu Leu Ser Glu Val Glu
                350
                                     355
Lys Ala Lys Ile Arg Gly Ser Tyr Leu Thr Val Thr Leu Gln Arg
                365
                                     370
Pro Thr Lys Glu Leu His Gly Thr Ser Ile Val Pro Lys Leu Gln
                380
                                     385
Ala Ile Thr Ala Ser Ser Ala Asn Leu Arg His Ser Pro Arg Val
                395
                                     400
Leu Val Gln His Cys Pro Ala Arg Thr Pro Gln Arg Gly Asp Glu
                 410
                                     415
Glu Gly Leu Gly Gly Ser Arg Arg Arg Lys Arg Arg Arg Arg Asp
                425
                                     430
Gly Gly Arg
<210> 183
<211> 246
<212> PRT
<213> Homo sapiens
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<400> 183
Gln Ala Leu Met Leu Val Ser Gly Arg Arg Arg Leu Leu Thr Ala
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Leu Leu Gln Ala Gln Lys Trp Pro Phe Gln Pro Ser Arg Asp Met
                 20
                                      25
Arg Leu Val Gln Phe Arg Ala Pro His Leu Val Gly Pro His Leu
                 35
                                      40
Gly Leu Glu Thr Gly Asn Gly Gly Gly Val Ile Asn Leu Asn Ala
                 50
                                      55
Phe Asp Pro Thr Leu Pro Lys Thr Met Thr Gln Phe Leu Glu Gln
                 65
                                     70
Gly Glu Ala Thr Leu Ser Val Ala Arg Arg Ala Leu Ala Ala Gln
                 80
                                      85
Leu Pro Val Leu Pro Trp Ser Glu Val Thr Phe Leu Ala Pro Val
                 95
                                     100
Thr Trp Pro Asp Lys Val Val Cys Val Gly Met Asn Tyr Val Asp
                110
                                     115
His Cys Lys Glu Gln Asn Val Pro Val Pro Lys Glu Pro Ile Ile
                125
                                    130
Phe Ser Lys Phe Ala Ser Ser Ile Val Gly Pro Tyr Asp Glu Val
                140
                                    145
Val Leu Pro Pro Gln Ser Gln Glu Val Asp Trp Glu Val Glu Leu
                155
                                     160
Ala Val Val Ile Gly Lys Lys Gly Lys His Ile Lys Ala Thr Asp
                170
                                    175
                                                         180
Ala Met Ala His Val Ala Gly Phe Thr Val Ala His Asp Val Ser
                185
                                    190
Ala Arg Asp Trp Leu Thr Arg Arg Asn Gly Lys Gln Trp Leu Leu
                200
                                    205
```

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Gly Lys Thr Phe Asp Thr Phe Cys Pro Leu Gly Pro Ala Leu Val
                215
                                    220
Thr Lys Asp Ser Val Ala Gly Arg Ser Leu Val Pro Ala Pro Trp
                                                         240
                                    235
                230
Tyr Leu Pro Leu His Arg
                245
<210> 184
<211> 266
<212> PRT
<213> Homo sapiens
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<223> Incyte ID No: LG:457464.24.orf3:2002JAN18
<400> 184
Gln Phe Ser Glu Gln Gln Pro Ser Tyr Gly Gly Gln Gln Ser
Tyr Gly Gln Gln Gln Ser Tyr Asn Pro Pro Gln Gly Tyr Gly Gln
                                                          30
                 20
                                      25
Gln Asn Gln Tyr Asn Ser Ser Ser Gly Gly Gly Gly Gly Gly
                                                          45
                                      40
                 35
Gly Gly Gly Asn Tyr Gly Gln Asp Gln Ser Ser Met Ser Ser Gly
                                                          60
                 50
                                      55
Gly Gly Ser Gly Gly Gly Tyr Gly Asn Gln Asp Gln Ser Gly Gly
                                      70
                 65
Gly Gly Ser Gly Gly Tyr Gly Gln Gln Asp Arg Gly Gly Arg Gly
                                                          90
                                      85
                 80
Arg Gly Gly Ser Gly Gly Ala Ala Ala Ala Ala Val Val Thr
                                     100
Thr Ala Ala Val Val Ala Met Asn Pro Glu Val Val Glu Val Ala
                                     115
                110
Val Glu Ala Glu Val Ala Trp Gly Pro Arg Asp Gln Gly Ser Arg
                                     130
                                                          135
                125
His Asp Ser Glu Gln Asp Asn Ser Asp Asn Asn Thr Ile Phe Val
                                     145
                                                          150
                140
Gln Gly Leu Gly Glu Asn Val Thr Ile Glu Ser Val Ala Asp Tyr
                                                          165
                155
                                     160
Phe Lys Gln Ile Gly Ile Ile Lys Thr Asn Lys Lys Thr Gly Gln
                                                          180
                 170
                                     175
Pro Met Ile Asn Leu Tyr Thr Asp Arg Glu Thr Gly Lys Leu Lys
                                     190
                 185
Gly Glu Ala Thr Val Ser Phe Asp Asp Pro Pro Ser Ala Lys Ala
                                     205
                                                          210
                 200
Ala Ile Asp Trp Phe Asp Gly Lys Glu Phe Ser Gly Asn Pro Ile
                 215
                                     220
Lys Val Ser Phe Ala Thr Arg Arg Ala Asp Phe Asn Arg Gly Gly
                                     235
                 230
Gly Asn Gly Arg Gly Gly Arg Gly Arg Gly Gly Pro Met Gly Arg
                 245
                                     250
Gly Gly Tyr Gly Gly Gly Ser Ala Gly Trp
                 260
                                     265
<210> 185
<211> 539
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<213> Homo sapiens
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<400> 185 Lys Leu Gly Ser Met Glu Pro Ala Pro Ala Arg Ser Pro Arg Pro Gln Gln Asp Pro Ala Arg Pro Gln Glu Pro Thr Met Pro Pro Glu Thr Pro Ser Glu Gly Arg Gln Pro Ser Pro Ser Pro Ser Pro Thr Glu Arg Ala Pro Ala Ser Glu Glu Glu Phe Gln Phe Leu Arg Cys Gln Gln Cys Gln Ala Glu Ala Lys Cys Pro Lys Leu Leu Pro Cys Leu His Thr Leu Cys Ser Gly Cys Leu Glu Ala Ser Gly Met Gln Cys Pro Ile Cys Gln Ala Pro Trp Pro Leu Gly Ala Asp Thr Pro Ala Leu Asp Asn Val Phe Phe Glu Ser Leu Gln Arg Arg Leu Ser Val Tyr Arg Gln Ile Val Asp Ala Gln Ala Val Cys Thr Arg Cys Lys Glu Ser Ala Asp Phe Trp Cys Phe Glu Cys Glu Gln Leu Leu Cys Ala Lys Cys Phe Glu Ala His Gln Trp Phe Leu Lys His Glu Ala Arg Pro Leu Ala Glu Leu Arg Asn Gln Ser Val Arg Glu Phe Leu Asp Gly Thr Arg Lys Thr Asn Asn Ile Phe Cys Ser Asn Pro Asn His Arg Thr Pro Thr Leu Thr Ser Ile Tyr Cys Arg Gly Cys Ser Lys Pro Leu Cys Cys Ser Cys Ala Leu Leu Asp Ser Ser His Ser Glu Leu Lys Cys Asp Ile Ser Ala Glu Ile Gln Gln Arg Gln Glu Glu Leu Asp Ala Met Thr Gln Ala Leu Gln Glu Gln Asp Ser Ala Phe Gly Ala Val His Ala Gln Met His Ala Ala Val Gly Gln Leu Gly Arg Ala Arg Ala Glu Thr Glu Glu Leu Ile Arg Glu Arg Val Arg Gln Val Val Ala His Val Arg Ala Gln Glu Arg Glu Leu Leu Glu Ala Val Asp Ala Arg Tyr Gln Arg Asp Tyr Glu Glu Met Ala Ser Arg Leu Gly Arg Leu Asp Ala Val Leu Gln Arg Ile Arg Thr Gly Ser Ala Leu Val Gln Arg Met Lys Cys Tyr Ala Ser Asp Gln Glu Val Leu Asp Met His Gly Phe Leu Arg Gln Ala Leu Cys Arg Leu Arg Gln Glu Glu Pro Gln Ser Leu Gln Ala Ala Val Arg Thr Asp Gly Phe Asp Glu Phe Lys Val Arg Leu Gln Asp Leu Ser Ser Cys Ile Thr Gln Gly Lys Asp Ala Ala Val Ser Lys Lys Ala Ser Pro Glu Ala Ala Ser Thr Pro Arg Asp Pro Ile Asp Val Asp Leu Ala Glu Glu Ala Glu Arg Val Lys Ala Gln Val Gln Ala Leu Gly Leu Ala Glu Ala Gln Pro Met Ala Val Val Gln Ser Val Pro Gly Ala His Pro Val Pro Val Tyr Ala Phe Ser Ile Lys Gly 

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Pro Ser Tyr Gly Glu Asp Val Ser Asn Thr Thr Thr Ala Gln Lys
                470
                                     475
Arg Lys Cys Ser Gln Thr Gln Cys Pro Arg Lys Val Ile Lys Met
                                                         495
                485
                                     490
Glu Ser Glu Glu Gly Lys Glu Ala Arg Leu Ala Arg Ser Ser Pro
                                     505
                500
Glu Gln Pro Arg Pro Ser Thr Ser Lys Ala Val Ser Pro Pro His
                515
                                     520
Leu Asp Gly Pro Pro Ser Pro Arg Ser Pro Val Ile Gly Ser
                530
                                   . 535
<210> 186
<211> 242
<212> PRT
<213> Homo sapiens
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Thr Gln Leu Thr Thr Asn Gln Thr Asn Pro Ser Gly Gln Ile Ser
                                      10
Tyr Glu Cys Gly Gln Cys Gly Arg Tyr Phe Ile Gln Met Ala Asp
                 20
                                      25
Phe His Arg His Glu Lys Cys His Thr Gly Glu Lys Ser Phe Glu
                 35
                                      40
Cys Lys Glu Cys Gly Lys Tyr Phe Arg Tyr Asn Ser Leu Leu Ile
                                                           60
                 50
Arg His Gln Ile Ile His Thr Gly Lys Lys Pro Phe Lys Cys Lys
                                      70
Glu Cys Gly Lys Gly Leu Ser Ser Asp Thr Ala Leu Ile Gln His
                                      85
                  80
Gln Arg Ile His Thr Gly Glu Lys Pro Tyr Glu Cys Lys Glu Cys
                                     100
                 95
Gly Lys Ala Phe Ser Ser Ser Val Phe Leu Gln His Gln Arg
                                     115
                                                          120
                110
Phe His Thr Gly Glu Lys Leu Tyr Glu Cys Asn Glu Cys Trp Lys
                                     130
                125
Thr Phe Ser Cys Ser Ser Ser Phe Thr Val His Gln Arg Met His
                 140
                                     145
                                                          150
Thr Gly Glu Lys Pro Tyr Glu Cys Lys Glu Cys Gly Lys Arg Leu
                                     160
                155
Ser Ser Asn Thr Ala Leu Thr Gln His Gln Arg Ile His Thr Gly
                                     175
                170
Glu Lys Pro Phe Glu Cys Lys Glu Cys Gly Lys Ala Phe Asn Gln
                 185
                                     190
                                                          195
Lys Ile Thr Leu Ile Gln His Gln Arg Val His Thr Gly Glu Lys
                                     205
                 200
Pro Tyr Glu Cys Lys Val Cys Gly Lys Thr Phe Ser Trp Cys Gly
                                                          225
                 215
                                     220
Arg Phe Ile Leu His Gln Lys Leu His Thr Gln Lys Thr Pro Val
                                     235
                 230
Gln Ala
<210> 187
<211> 194
<212> PRT
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<223> Incyte ID No: LG:7689661.4.orf2:2002JAN18 <400> 187 Asn Val Lys Glu Cys Gly Lys Ala Phe Arg Val Arg Gly Gln Leu Thr Leu His Gln Arg Ile His Thr Gly Glu Lys Pro Tyr Glu Cys 20 25 30 Lys Glu Cys Gly Lys Thr Phe Ser Arg Gly Tyr His Leu Ile Leu 40 His His Arg Ile His Thr Gly Glu Lys Pro Tyr Glu Cys Lys Glu 50 Cys Trp Lys Ala Phe Ser Arg Tyr Ser Gln Leu Ile Ser His Gln 65 70 Ser Ile His Ile Gly Val Lys Pro Tyr Asp Cys Lys Glu Cys Gly 80 85 Lys Ala Phe Arg Leu Leu Ser Gln Leu Thr Gln His Gln Ser Ile 95 100 His Ile Gly Glu Lys Pro Tyr Lys Cys Lys Glu Cys Gly Lys Ala 110 115 120 Phe Arg Leu Arg Gln Lys Leu Thr Leu His Gln Ser Ile His Thr 125 130 Gly Glu Lys Pro Phe Glu Cys Lys Glu Cys Arg Lys Ala Phe Arg 140 145 Leu Asn Ser Ser Leu Ile Gln His Leu Arg Ile His Ser Gly Glu 155 160 Lys Pro Tyr Glu Cys Lys Glu Cys Lys Lys Ala Phe Arg Gln His 170 175 Ser His Leu Thr His His Leu Lys Ile His Asn Val Lys Ile 185 190 <210> 188 <211> 149 <212> PRT <213> Homo sapiens <220> <221> misc_feature <223> Incyte ID No: LG:7690373.1.orf1:2002JAN18 <400> 188 Lys Asn Ser Tyr Trp Arg Lys Asn Pro Thr Asn Met Lys Asn Val 10 Ala Lys Leu Leu Ile Asn Ser Gln Arg Leu Leu Asn Ile Arg Glu 20 25 30 Phe Val Glu Glu Gly Asn Pro Thr Asn Leu Lys Asn Val Ala Ser 35 40 Leu Leu Ala Ile Pro Gln Ser Leu Leu Asn Ile His Val Ile His 50 55 Thr Gly Gly Asn Ser Tyr Asn Cys Val Giu Cys Cys Asn Ala Leu 65 Asn Gln Ser Leu Arg Leu Thr Thr Tyr Lys Thr Thr His Thr Gly 80 85 90 Glu Lys Pro Cys Met Cys Glu Glu Cys Gly Lys Ala Ser Asn Arg 95 100 Ser Ser Ile Leu Lys Arg His Lys Leu Ile His Thr Gln Glu Arg 110 115 Leu Tyr Lys Pro Glu Arg Cys Asp Asn Ala Phe Gly Asn Thr Ser 125 130 Asp Phe Ser Glu Tyr Lys Arg Asn Arg Thr Asp Glu Lys Ser 140 145 <210> 189 <211> 268

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<213> Homo sapiens
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Lys Val Ala Thr Met Arg Lys Leu Phe Ser Phe Gly Arg Arg Leu
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Gly Gln Ala Leu Leu Asp Ser Met Asp Gln Glu Tyr Ala Gly Arg
                                     25
                 20
Gly Tyr His Ile Arg Asp Trp Glu Leu Arg Lys Ile His Arg Ala
                                     40
                 35
Ala Ile Lys Gly Asp Ala Ala Glu Val Glu His Cys Leu Thr Arg
                 50
                                     55
Arg Phe Arg Asp Leu Asp Ala Arg Asp Arg Lys Asp Arg Thr Val
                                     70
                 65
Leu His Leu Thr Cys Ala His Gly Arg Val Glu Val Val Thr Leu
                                     85
                 80
Leu Leu Ser Arg Arg Cys Gln Ile Asn Ile Tyr Asp Arg Leu Asn
                 95
                                    100
Arg Thr Pro Leu Met Lys Ala Val His Cys Gln Glu Glu Ala Cys
                                                         120
                                    115
                110
Ala Ile Ile Leu Leu Glu His Gly Ala Asn Pro Asn Ile Lys Asp
                                                         135
                                    130
                125
Ile Tyr Ser Asn Thr Ala Leu His Tyr Ala Val Tyr Asn Lys Gly
                                    145
                140
Thr Ser Leu Ala Glu Lys Leu Leu Ser His His Ala Asn Ile Glu
                                     160
                155
Ala Leu Asn Glu Glu Gly Asn Thr Pro Leu Leu Phe Ala Ile Asn
                                     175
                170
Ser Arg Arg Gln Gln Ile Val Glu Phe Leu Leu Lys Asn Gln Ala
                                     190
                185
Asn Leu His Ala Ile Asp Asn Phe Arg Arg Thr Ala Leu Met Leu
                                                         210
                                     205
                200
Ala Val Gln His Asn Ser Ser Ser Ile Val Ser Leu Leu Gln
                                  220
                215
Gln Asn Ile Asn Ile Phe Ser Gln Asp Leu Phe Gly Gln Thr Ala
                                     235
                230
Glu Asp Tyr Ala Val Cys Tyr Asn Phe Arg Ser Ile Gln Gln
                                     250
                245
Ile Leu Glu His Lys Asn Lys Ile Leu Lys Ser His Leu
                                     265
                260
<210> 190
<211> 1304
<212> PRT
<213> Homo sapiens
<220>
<221> misc_feature
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<220>
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 <222> (1) ... (1304)
 <223> unknown or other
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 Arg Trp Arg Lys Leu Pro Lys Met Pro Glu Ala Val Gly Thr Asp
                                      10
 Pro Ser Thr Ser Arg Lys Met Ala Glu Leu Glu Glu Val Thr Leu
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			Pro	35					40		_		_	45
			Gln	50					55			_		60
			Arg	65					70					75
			Thr	80					85					90
			Met	95					100					105
			Glu	110					115					120
			Glu	125					130					135
			Pro	140					145				_	150
			Leu	155					160					165
	٠.		Ser -	170					175					180
			Pro	185					190					195
			Ala	200					205					210
			Asp	215					220					225
			Asn	230					235					240
			Thr Arg	245					250					255
			Met	260					265					270
			Thr	275					280					285
				290					295				_	300
			Thr	305					310					315
			Gln	320					325					330
			Glu	335	•				340					345
пув	ser	ьys	Ser	350	ser	Pro	Pro	Arg	ьеи 355	Thr	Glu	Asp	Arg	Lys 360
			Leu	365					370					375
			Pro	380					385					390
			Gln	395					400					405
			Pro	410					415					420
			Thr	425					430					435
			Gly	440					445					450
ATa	GIU	Ser	Pro	Ala 455	Glu	Lys	Val	Pro	Glu 460	Glu	Ser	Val	Leu	Pro 465
Leu	Val	Gln	Lys		Thr	Leu	Ala	Asp		Ser	Ala	Gln	Lys	
Leu	Glu	Pro	Glu	Ser 485	Asp	Arg	Ser	Ala		Pro	Leu	Pro	Leu	

	Ile	Glu	Glu	Leu	Ala 500	Leu	Ala	Lys	Gly	Ile 505	Thr	Glu	Glu	Cys	Leu 510
	Lys	Gln	Pro	Ser	Leu 515	Glu	Gln	Lys	Glu	Gly 520	Arg	Arg	Ala	Ser	His 525
	Thr	Leu	Leu	Pro	Ser 530	His	Arg	Leu	Lys	Gln 535	Ser	Ala	Asp	Ser	Ser 540
	Ser	Ser	Arg	Ser	Ser 545	Ser	Ser	Ser	Ser	Ser 550	Ser	Ser	Arg	Ser	Arg 555
		_			560					565				Pro	570
	Arg	Ser	Lys	Gln	Arg 575	Asp	Val	Ala	Gln	Ala 580	Arg	Thr	His	Ala	Asn 585
		_	_		590	_		_		595				Glu	600
	_				605					610				Asn	615
					620					625				Thr	630
					635					640				Ala	645
					650					655				Arg	660
					665					670				Gln	675
					680					685				Glu	690
					695					700				Pro	705
					710				•	715				Leu	720
					725					730				Glu	735
			_		740	•		_		745				Pro	750
					755					760				Arg	765
		• -			770	_				775				Leu	780
	_				785	_				790				Lys	795
					800					805				Gly	810
	_	_	_	_	815					820			_	Lys	825
					830					835					Asp 840
					845	-				850				His	855
					860			_		865				Gly Val	870
					875					880				Glu	885
					890		_			895				Pro	900
					905					910				His	915
					920					925				Ser	930
					935		_	_		940				Pro	945
					950					955				Ser	960
•	AT 9	T 11T	тта	GAII	val	F <b>T</b> O	Ser	-10	ETO	Arg	GTY	y S		CEL	

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965
                                     970
 Ile Val His Ile Ser Asn Leu Val Arg Pro Phe Thr Leu Gly Gln
                                     985
 Leu Lys Glu Leu Leu Gly Arg Thr Gly Thr Leu Val Glu Glu Ala
                 995
                                    1000
                                                        1005
 Phe Trp Ile Asp Lys Ile Lys Ser His Cys Phe Val Thr Tyr Ser
                1010
                                    1015
 Thr Val Glu Glu Ala Val Ala Thr Arg Thr Ala Leu His Gly Val
                1025
                                    1030
                                                        1035
. Lys Trp Pro Gln Ser Asn Pro Lys Phe Leu Cys Ala Asp Tyr Ala
                1040
                                    1045
                                                        1050
 Glu Gln Asp Glu Leu Asp Tyr His Arg Gly Leu Leu Val Asp Arg
                1055
                                    1060
 Pro Ser Glu Thr Lys Thr Glu Glu Gln Gly Ile Pro Arg Pro Leu
                1070
                                    1075
 His Pro Pro Pro Pro Pro Val Gln Pro Pro Gln His Pro Arg
                1085
                                    1090
Ala Glu Gln Arg Glu Gln Glu Arg Ala Val Arg Glu Gln Trp Ala
                1100
                                   1105
                                                        1110
Glu Arg Glu Arg Glu Met Glu Arg Arg Glu Arg Thr Arg Ser Glu
                1115
                                    1120
                                                        1125
Arg Glu Trp Asp Arg Asp Lys Val Arg Glu Gly Pro Arg Ser Arg
                1130
                                    1135
                                                        1140
Ser Arg Ser Arg Asp Arg Arg Lys Glu Arg Ala Lys Ser Lys
                1145
                                    1150
Glu Lys Lys Ser Glu Lys Lys Glu Lys Ala Gln Glu Glu Pro Pro
                1160
                                   1165
Ala Lys Leu Leu Asp Asp Leu Phe Arg Lys Thr Lys Ala Ala Pro
               1175
                                    1180
                                                        1185
Cys Ile Tyr Trp Leu Pro Leu Thr Asp Ser Gln Ile Val Gln Lys
               1190
                                   1195
                                                        1200
Glu Ala Glu Arg Ala Glu Arg Ala Lys Glu Arg Glu Lys Arg Arg
               1205
                                   1210
Lys Glu Glu Glu Glu Glu Lys Glu Arg Glu Lys Glu Ala
               1220
                                   1225
Glu Arg Glu Arg Asn Arg Gln Leu Glu Arg Glu Lys Arg Arg Glu
               1235
                                   1240
His Ser Arg Glu Arg Asp Arg Xaa Arg Xaa Arg Glu Arg Glu Arg
               1250
                                   1255
Asp Arg Gly Asp Arg Asp Arg Glu Arg Asp Arg Glu Arg
               1265
                                   1270
Gly Arg Glu Arg Asp Arg Asp Thr Lys Arg His Ser Arg Ser
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Arg Ser Arg Ser Thr Pro Val Arg Asp Arg Gly Gly Arg Arg
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Ser Arg Cys Xaa Val Thr Arg Gly Ser Gln Ala Trp Leu Pro Leu
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Leu Arg Pro Met Val Ala Ala Arg Trp Gly Ala Thr Val Gly Pro
Gly Ala Val Trp Thr Gln Cys Tyr Gly Trp Gly Trp Pro Glu Pro
                 50
                                     55
Ala Trp Asp Ser Arg Glu Trp Arg Arg Val Val Gly Pro Gly Lys
                                     70
Arg Pro Arg Leu Leu Ala His Pro Leu Trp Ala Ser Leu Glu Leu
                                     85
Leu Phe Leu Val Ser Gln Glu Asp Thr Leu Ser Pro Gly Ala Val
                                    100
                 95
Gly Pro Arg Cys Val Gly Asp Pro Gly Ser Ala Leu Gly Pro Leu
                110
                                    115
                                                         120
His Val Gly Asp Thr Gly Asn Ala Arg Ser Pro Pro Cys Phe Ser
                125
                                    130
Pro His Leu Pro Ile Ser Thr Cys Gly Ala Arg Gly Ser Asp Pro
                140
                                    145
Lys Ala Ala Ser His Pro Pro Ser Pro Ala Pro Pro Ala Leu Arg
                155
                                    160
Ala Gln Gly Ala Ala Gln Pro Cys His Leu Cys Ser Ser Pro Ala
                170
                                    175
Pro His Thr Asn Leu Gly Pro Gly Gly Pro Ala His Pro Gly Leu
                                     190
                185
Arg Arg Pro Pro Pro Leu Val His Met Ala Ser Pro Ser Cys Arg
                                                         210
                200
Gly Ser Gly Cys Cys Pro His Arg Ala Gly Ser Leu Leu Arg Cys
                                    220
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Ala Gly Lys Ala Gly Trp Cys Arg Gly Ala Arg Arg Gly Arg
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Pro Ala Leu Leu Ser Leu Val Leu Pro Ser Gln Gly Glu Ala Pro
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Ala Glu Met Gly Ala Leu Leu Glu Lys Glu Thr Arg Gly Ala
                                      25
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Thr Glu Arg Val His Gly Ser Leu Gly Asp Thr Pro Arg Ser Glu
Glu Thr Leu Pro Lys Ala Thr Pro Asp Ser Leu Glu Pro Ala Gly
                 50
                                      55
Pro Ser Ser Pro Ala Ser Val Thr Val Thr Val Gly Asp Glu Gly
                 65
                                      70
Ala Asp Thr Pro Val Gly Ala Thr Pro Leu Ile Gly Asp Glu Ser
                 80
                                      85
Glu Asn Leu Glu Gly Asp Gly Asp Leu Arg Gly Gly Arg Ile Leu
                 95
                                     100
                                                         105
Leu Gly His Ala Thr Lys Ser Phe Pro Ser Ser Pro Ser Lys Gly
                110
                                     115
                                                         120
Gly Ser Cys Pro Ser Arg Ala Lys Met Ser Met Thr Gly Ala Gly
                                     130
                                                         135
                125
Lys Ser Pro Pro Ser Val Gln Ser Leu Ala Met Arg Leu Leu Ser
                140
                                     145
                                                         150
Met Pro Gly Ala Gln Gly Ala Ala Ala Ala Gly Ser Glu Pro Pro
                                     160
                                                         165
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Pro	Ala	a Th:	r Th	r Se:	r Pro	o Glu	ı Gly	y Glr	n Pro 175	Lys	val	. His	arç	Ala
Arg	J Lys	s Thi	r Me	t Se: 18:	r Ly:	s Pro	Gly	/ Asr	1 Gly 190	/ Glr	Pro	Pro	Va.	180 L Pro
Gli	ı Lys	a Arg	g Pro	200	Glı	ı Ile	e Glr	n His	Phe 205	Arg	, Met	Ser	Ası	195 Asp
Va]	His	S Sea	r Let	ي Gl 21	y Lys	s Val	Thi	Ser	203 Asp 220	Let	Ala	Lys	Arç	210 Arg
Lys	Lei	ı Ası	n Sei	c Gly 230	y Gly	/ Gly	/ Let	. Ser	Glu 235	Glu	Leu	Gly	Ser	225 Ala
Arc	J Arc	Sei	c Gly	/ Glu 245	ı Val	l Thr	Let	Thr	Lys 250	Gly	Asp	Pro	Gly	240 Ser
Leu	Glu	ı Glı	ı Trı	Glu 260	ı Thi	. Val	. Val	. Gly	Asp 265	Asp	Phe	Ser	Lev	255 Tyr 270
				275	)				Val 280	Asp				Lys
				290	)				Leu 295	Ser				Glu
				305	)				Glu 310	Glu				Glu
				320	)				325					Arg
				333	)				340					Lys
				350	)				355					Glu
				365	,	Pro			370					Ser
				380		Gly			385					Ser
				395		Lys			400					405
				4±0		Thr			415					120
				425		Leu			430					125
	•			440		Val			445					450
				455		Суз			460					165
				4/0		Сув			475					10n
				485		Val			4 D D					40-
				200		Asp			ついつ					E10
				272		Val			520					525
				230		Ser			535					540
				245		Arg			550					555
				טסכ		Leu Asn			565					ETA
				3/5		Arg			580					505
				<b>590</b>		Arg			595					600
				605		Lys			610					615
				020		Leu			625					630
							-2010	0	U911	-IIG	GIII	ser .	ASP	GIII

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640
                635
Gln Ser Lys Arg Thr Pro Leu His Ala Ala Ala Gln Lys Gly Ser
                650
                                     655
Val Glu Ile Cys His Val Leu Leu Gln Ala Gly Ala Asn Ile Asn
                                     670
                665
Ala Val Asp Lys Gln Gln Arg Thr Pro Leu Met Glu Ala Val Val
                                     685
                680
Asn Asn His Leu Glu Val Ala Arg Tyr Met Val Gln Arg Gly Gly
                695
                                     700
Cys Val Tyr Ser Lys Glu Glu Asp Gly Ser Thr Cys Leu His His
                710
                                     715
Ala Ala Lys Ile Gly Asn Leu Glu Met Val Ser Leu Leu Leu Ser
                                                         735
                725
                                     730
Thr Gly Gln Val Asp Val Asn Ala Gln Asp Ser Gly Gly Trp Thr
                740
                                     745
Pro Ile Ile Trp Ala Ala Glu His Lys His Ile Glu Val Ile Arg
                755
                                     760
Met Leu Leu Thr Arg Gly Ala Asp Val Thr Leu Thr Asp Asn Val
                                     775
                                                         780
                770
Ser Glu Arg Leu Val Glu Val Gly Gln Pro Gln Ala Pro Glu Gln
                785
                                     790
Gly Gly Gly Trp Ile Gln Gly Pro Ser Cys Cys Thr Ser Ser Val
                                                      810
                800
                                     805
Pro Leu Leu Pro Pro Gln Glu Glu Asn Ile Cys Leu His Trp Ala
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                815
Ser Phe Thr Gly Ser Ala Ala Ile Ala Glu Val Leu
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Arg Leu Ile Asn Gly Leu Gly Cys Lys Leu Ser Phe Ile Pro Trp
                                      25
                                                          30
                  20
Asp Ala Leu Ser Ala Leu Gln His Leu Lys Phe Arg Gln Arg Glu
                 35
                                      40
                                                          45
Leu Thr Trp Gly Gln Ala Ala Pro Leu Gly Arg Val Glu Asp Arg
                                      55
                 50
Val Ser Leu Leu Ile Phe Arg Lys Ser Ser Arg Thr Gln Ser Pro
                                      70
Ala Phe Gly Ser Leu Ser Gln Arg Asp Arg Arg Asn Pro Glu Gln
                                      85
                  80
Ala Thr Gly Arg Arg Ser Gly Met Tyr Phe Cys Trp Gly Ala Asp
                                     100
                  95
Ser Arg Glu Leu Gln Arg Arg Thr Ala Gly Ser Pro Gly Ala
                 110
                                     115
Glu Leu Leu Gln Ala Ala Ser Gly Glu Arg His Ser Leu Leu
                                     130
                                                         135
                125
Leu Thr Asn His Arg Val Leu Ser Cys Gly Asp Asn Ser Arg Gly
                140
                                     145
                                                          150
Gln Leu Gly Arg Arg Gly Ala Gln Arg Gly Glu Leu Pro Glu Pro
                                     160
                 155
Ile Gln Ala Leu Glu Thr Leu Ile Val Asp Leu Val Ser Cys Gly
                                     175
                                                          180
                 170
Lys Glu His Ser Leu Ala Val Cys His Lys Gly Arg Val Phe Ala
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185
                                      190
 Trp Gly Ala Gly Ser Glu Gly Gln Leu Gly Ile Gly Glu Phe Lys
                 200
                                      205
                                                          210
Glu Ile Ser Phe Thr Pro Lys Lys Ile Met Thr Leu Asn Asp Ile
                 215
                                      220
                                                           225
 Lys Ile Ile Gln Val Ser Cys Gly His Tyr His Ser Leu Ala Leu
                 230
                                      235
 Ser Lys Asp Ser Gln Val Phe Ser Trp Gly Lys Asn Ser His Gly
                 245
                                      250
                                                          255
Gln Leu Gly Leu Gly Lys Glu Phe Pro Ser Gln Ala Ser Pro Gln
                 260
                                     265
Arg Val Arg Ser Leu Glu Gly Ile Pro Leu Ala Gln Val Ala Ala
                 275
                                     280
Gly Gly Ala His Ser Phe Ala Leu Ser Leu Cys Gly Thr Ser Phe
                 290
                                     295
Gly Trp Gly Ser Asn Ser Ala Gly Gln Leu Ala Leu Ser Gly Arg
                 305
                                     310
Asn Val Pro Val Gln Ser Asn Lys Pro Leu Ser Val Gly Ala Leu
                 320
                                     325
                                                          330
Lys Asn Leu Gly Val Val Tyr Ile Ser Cys Gly Asp Ala His Thr
                 335
                                     340
                                                          345
Ala Val Leu Thr Gln Asp Gly Lys Val Phe Thr Phe Gly Asp Asn
                 350
                                     355
                                                          360
Arg Ser Gly Gln Leu Gly Tyr Ser Pro Thr Pro Glu Lys Arg Gly
                 365
                                     370
                                                          375
Pro Gln Leu Val Glu Arg Ile Asp Gly Leu Val Ser Gln Ile Asp
                 380
                                     385
                                                          390
Cys Gly Ser Tyr His Thr Leu Ala Tyr Val His Thr Thr Gly Gln
                 395
                                     400
                                                          405
Val Val Ser Phe Gly His Gly Pro Ser Asp Thr Ser Lys Pro Thr
                 410
                                     415
                                                          420
His Pro Glu Ala Leu Thr Glu Asn Phe Asp Ile Ser Cys Leu Ile
                 425
                                     430
                                                          435
Ser Ala Glu Glu Thr Leu Ser Met Asp Leu
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Ile Thr Leu Leu Arg His Leu Asn Thr Ser Ser Leu Leu Cys Asp
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                                      25
Cys Gln Leu Lys Trp Leu Pro Gln Trp Val Ala Glu Asn Asn Phe
                 35
                                      40
Gln Ser Phe Val Asn Ala Ser Cys Ala His Pro Gln Leu Leu Lys
                 50
                                      55
Gly Arg Ser Ile Phe Ala Val Ser Pro Asp Gly Phe Val Cys Asp
                 65
                                      70
Asp Phe Pro Lys Pro Gln Ile Thr Val Gln Pro Glu Thr Gln Ser
                 80
                                      85
Ala Ile Lys Gly Ser Asn Leu Ser Phe Ile Cys Ser Ala Ala Ser
                 95
                                     100
Ser Ser Asp Ser Pro Met Thr Leu Leu Gly Lys Lys Thr Met Asn
                110
                                     115
Tyr Cys Met Met Leu Lys Trp Lys Ile Met His Thr Ser Gly Pro
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125 130 135

Lys Val Ala Arg

<210> 195

<211> 650

<212> PRT

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<400> 195

Leu Pro Phe Ser Glu Asp Gly Ser Ser Val Pro His Ile Cys His 10 Val His Pro Gly Phe His Leu Ser Pro Gly Leu Arg Ile Ser Cys 30 20 Phe Phe Lys Arg Pro Phe Leu Ser Pro Glu Phe Gly Pro Val Arg 45 40 35 Val Gln Trp Ser Gly Ala Ser His Thr Gln Cys Trp Phe Pro Gly 50 55 Ile Gly Asp Phe Pro Arg Cys Arg Cys Gly Leu Tyr Arg Glu Gly 70 65 Val Ala Leu Ala Gly Phe Phe Ser Glu Lys Thr Val Gln Arg Cys 90 80 85 Asn Ala Gly Glu Leu Gln Gln Pro His Phe Thr Gly Asn Phe Gly 95 100 Thr Thr His Phe Ala Ala Pro Lys Ser Asp Leu Ser Thr Leu Arg 115 120 110 Ser Ile Glu Asp Pro Ser Val Glu Pro Arg Leu Leu Glu Gly Val 130 125 Val Pro Leu His Gly Pro Pro Ser Thr Cys Val Phe Pro Val Ser 145 140 Val Gly Tyr Gln Val Gly Lys Pro Ser Leu Ile Ser His Leu Glu 155 160 Gln Glu Glu Pro Arg Thr Glu Glu Arg Gly Ala His Gln Gly 175 170 Ala Cys Ala Asp Trp Glu Thr Pro Ser Lys Thr Lys Trp Ser Leu 190 195 185 Leu Met Glu Asp Ile Phe Gly Lys Glu Thr Pro Ser Gly Val Thr 210 205 200 Met Glu Arg Ala Gly Leu Gly Glu Lys Ser Thr Glu Tyr Ala His 220 215 Leu Phe Glu Val Phe Gly Met Asp Pro His Leu Thr Gln Pro Met 230 235 240 Gly Arg His Ala Gly Lys Arg Pro Tyr His Arg Arg Asp Tyr Gly 255 245 250 Val Ala Phe Lys Gly Arg Pro His Leu Thr Gln His Met Ser Met 260 270 Tyr Asp Gly Arg Lys Met His Glu Cys His Gln Cys Gln Lys Ala 280 275 Phe Thr Thr Ser Ala Ser Leu Thr Arg His Arg Arg Ile His Thr 295 290 Gly Glu Lys Pro Tyr Glu Cys Ser Asp Cys Gly Lys Ala Phe Asn 305 310 Asp Pro Ser Ala Leu Arg Ser His Ala Arg Thr His Leu Lys Glu 325 330 320 Lys Pro Phe Asp Cys Ser Gln Cys Gly Asn Ala Phe Arg Thr Leu 340 335 Ser Ala Leu Lys Ile His Met Arg Val His Thr Gly Glu Arg Pro 350 355 Tyr Lys Cys Asp Gln Cys Gly Lys Ala Tyr Gly Arg Ser Cys His

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365
                                      370
                                                          375
Leu Ile Ala His Lys Arg Thr His Thr Gly Glu Arg Pro Tyr Glu
                 380
                                      385
                                                          390
Cys His Asp Cys Gly Lys Ala Phe Gln His Pro Ser His Leu Lys
                 395
                                      400
                                                          405
Glu His Val Arg Asn His Thr Gly Glu Lys Pro Tyr Ala Cys Thr
                 410
                                      415
Gln Cys Gly Lys Ala Phe Arg Trp Lys Ser Asn Phe Asn Leu His
                 425
                                     430
Lys Lys Asn His Met Val Glu Lys Thr Tyr Glu Cys Lys Glu Cys
                 440
                                      445
Gly Lys Ser Phe Gly Asp Leu Val Ser Arg Arg Lys His Met Arg
                 455
                                     460
Ile His Ile Val Lys Lys Pro Val Glu Cys Arg Gln Cys Gly Lys
                 470
                                      475
Thr Phe Arg Asn Gln Ser Ile Leu Lys Thr His Met Asn Ser His
                 485
                                     490
                                                          495
Thr Gly Glu Lys Pro Tyr Gly Cys Asp Leu Cys Gly Lys Ala Phe
                 500
                                     505
Ser Ala Ser Ser Asn Leu Thr Ala His Arg Lys Ile His Thr Gln
                 515
                                     520
                                                          525
Glu Arg Arg Tyr Glu Cys Ala Ala Cys Gly Lys Val Phe Gly Asp
                 530
                                     535
                                                          540
Tyr Leu Ser Arg Arg Arg His Met Ser Val His Leu Val Lys Lys
                 545
                                     550
                                                          555
Arg Val Glu Cys Arg Gln Cys Gly Lys Ala Phe Arg Asn Gln Ser
                560
                                     565
                                                          570
Thr Leu Lys Thr His Met Arg Ser His Thr Gly Glu Lys Pro Tyr
                575
                                     580
                                                          585
Glu Cys Asp His Cys Gly Lys Ala Phe Ser Ile Gly Ser Asn Leu
                590
                                     595
Asn Val His Arg Arg Ile His Thr Gly Glu Lys Pro Tyr Glu Cys
                 605
                                     610
Leu Val Cys Gly Lys Ala Phe Ser Asp His Ser Ser Leu Arg Ser
                620
                                     625
His Val Lys Thr His Arg Gly Glu Lys Leu Phe Val Ser Ser Val
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Trp Lys Arg Leu Gln
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Met Ala Met Ala Leu Pro Met Pro Gly Pro Gln Glu Ala Val Val
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Phe Glu Asp Val Ala Val Tyr Phe Thr Arg Ile Glu Trp Ser Cys
                 35
                                      40
Leu Ala Pro Asp Gln Gln Ala Leu Tyr Arg Asp Val Met Leu Glu
                 50
                                      55
Asn Tyr Gly Asn Leu Ala Ser Leu Gly Phe Leu Val Ala Lys Pro
                 65
                                      70
Ala Leu Ile Ser Leu Leu Glu Gln Gly Glu Glu Pro Gly Ala Leu
                                      85
Ile Leu Gln Val Ala Glu Gln Ser Val Ala Lys Ala Ser Leu Cys
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95
                                    100
                                                         105
Thr Glu Asp Pro Asn Thr Leu Pro Ser Arg Ser Gln Glu Gly Ser
                                    115
                110
Pro Ala Ser Ser Glu Gly Gly Pro Gly Glu Lys Gly Val Ala Gly
                                    130
                125
Arg Val Ala Gly Gly Gly Ala Ala Ser Ser Trp Pro His Gly Glu
                140
                                    145
His Pro Val Thr Pro Asn Arg
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Arg Leu Trp Leu Lys Phe His Arg His Gln Thr Glu Met Ile Ile
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Thr Lys Gln Gly Arg Arg Met Phe Pro Phe Leu Ser Phe Asn Ile
                 20
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Asn Gly Leu Asn Pro Thr Ala His Tyr Asn Val Phe Val Glu Val
                 35
                                     40
Val Leu Ala Asp Pro Asn His Trp Arg Phe Gln Gly Gly Lys Trp
                 50
                                      55
Val Thr Cys Gly Lys Ala Asp Asn Asn Met Gln Gly Asn Lys Met
                                     70
                 65
Tyr Val His Pro Glu Ser Pro Asn Thr Gly Ser His Trp Met Arg
                                      85
                 80
Gln Glu Ile Ser Phe Gly Lys Leu Lys Leu Thr Asn Asn Lys Gly
                                    100
                 95
Ala Asn Asn Asn Thr Gln Met Ile Val Leu Gln Ser Leu His
                110
                                    115
                                                      120
Lys Tyr Gln Pro Arg Leu His Ile Val Glu Val Thr Glu Asp Gly
                125
                                     130
                                                         135
Val Glu Asp Leu Asn Glu Pro Ser Lys Thr Gln Thr Phe Thr Phe
                140
                                                         150
                                     145
Ser Glu Thr Gln Phe Ile Ala Val Thr Ala Tyr Gln Asn Thr Asp
                155
                                     160
Ile Thr Gln Leu Lys Ile Asp His Asn Pro Phe Ala Lys Gly Phe
                170
                                     175
                                                         180
Arg Asp Asn Tyr Asp Ser Met Tyr Thr Ala Ser Glu Asn Asp Arg
                                                         195
                185
                                     190
Leu Thr Pro Ser Pro Thr Asp Ser Pro Arg Ser His Gln Ile Val
                                     205
                200
Pro Gly Gly Arg Tyr Gly Val Gln Ser Phe Phe Pro Glu Pro Phe
                                     220
                                                         225
                215
Val Asn Thr Leu Pro Gln Ala Arg Tyr Tyr Asn Gly Glu Arg Thr
                                     235
                230
Val Pro Gln Thr Asn Gly Leu Leu Ser Pro Gln Gln Ser Glu Glu
                                                         255
                245
                                     250
Val Ala Asn Pro Pro Gln Arg Trp Leu Val Thr Pro Val Gln Gln
                260
                                     265
                                                         270
Pro Gly Thr Asn Lys Leu Asp Ile Ser Ser Tyr Glu Ser Glu Tyr
                275
                                     280
                                                         285
Thr Ser Ser Thr Leu Leu Pro Tyr Gly Ile Lys Ser Leu Pro Leu
                290
                                     295
Gln Thr Ser His Ala Leu Gly Tyr Tyr Pro Asp Pro Thr Phe Pro
                305
                                     310
Ala Met Ala Gly Trp Gly Gly Arg Gly Ser Tyr Gln Arg Lys Met
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320
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Ala Ala Gly Leu Pro Trp Thr Ser Arg Thr Ser Pro Thr Val Phe
                 335
                                     340
Ser Glu Asp Gln Leu Ser Lys Glu Lys Val Lys Glu Glu Ile Gly
                 350
                                     355
                                                          360
Ser Ser Trp Ile Glu Thr Pro Pro Ser Ile Lys Ser Leu Asp Ser
                 365
                                     370
Asn Asp Ser Gly Val Tyr Thr Ser Ala Cys Lys Arg Arg Leu
                 380
                                     385
Ser Pro Ser Asn Ser Ser Asn Glu Asn Ser Pro Ser Ile Lys Cys
                 395
                                     400
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Glu Asp Ile Asn Ala Glu Glu Tyr Ser Lys Asp Thr Ser Lys Gly
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                                     415
Met Gly Gly Tyr Tyr Ala Phe Tyr Thr Thr Pro
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Gln Arg Ile Arg Ala Ala Asn Ser Asn Gly Leu Pro Arg Cys Lys
                 20
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Ser Glu Gly Thr Leu Ile Asp Leu Ser Glu Gly Phe Ser Glu Thr
                 35
                                      40
Ser Phe Asn Asp Ile Lys Val Pro Ser Pro Ser Ala Leu Leu Val
                 50
                                      55
Asp Asn Pro Thr Pro Phe Gly Asn Ala Lys Glu Val Ile Ala Ile
                 65
                                      70
Lys Asp Tyr Cys Pro Thr Asn Phe Thr Thr Leu Lys Phe Ser Lys
                 80
                                      85
Gly Asp His Leu Tyr Val Leu Asp Thr Ser Gly Gly Glu Trp Trp
                 95
                                     100
                                                         105
Tyr Ala His Asn Thr Thr Glu Met Gly Tyr Ile Pro Ser Ser Tyr
                110
                                     115
                                                         120
Val Gln Pro Leu Asn Tyr Arg Asn Ser Thr Leu Ser Asp Ser Gly
                125
                                     130
                                                         135
Met Ile Asp Asn Leu Pro Asp Ser Pro Asp Glu Val Ala Lys Glu
                140
                                     145
Leu Glu Leu Leu Gly Gly Trp Thr Asp Asp Lys Lys Val Pro Gly
                155
                                     160
Arg Met Tyr Ser Asn Asn Pro Phe Trp Asn Gly Val Gln Thr Asn
                170
                                     175
Pro Phe Leu Asn Gly Asn Val Pro Val Met Pro Ser Leu Asp Glu
                185
                                     190
Leu Asn Pro Lys Ser Thr Val Asp Leu Leu Leu Phe Asp Ala Gly
                200
                                     205
Thr Ser Ser Phe Thr Glu Ser Ser Ser Ala Thr Thr Asn Ser Thr
                215
                                     220
Gly Asn Ile Phe Asp Glu Leu Pro Val Thr Asn Gly Leu His Ala
                230
                                     235
Glu Pro Pro Val Arg Arg Asp Asn Pro Phe Phe Arg Ser Lys Arg
                245
                                     250
Ser Tyr Ser Leu Ser Glu Leu Ser Val Leu Gln Ala Lys Ser Asp
                260
                                     265
Ala Pro Thr Ser Ser Ser Phe Phe Thr Gly Leu Lys Ser Pro Ala
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				275					280					285
Pro	Glu	Gln	Phe	Gln 290	Ser	Arg	Glu	Asp	Phe 295	Arg	Thr	Ala	Trp	Leu 300
Asn	His	Arg	Lys	Leu 305	Ala	Arg	Ser	Cys	His 310	Asp	Leu	Asp	Leu	Leu 315
Gly	Gln	Ser	Pro	Gly 320	Trp	Gly	Gln	Thr	Gln 325	Ala	Val	Glu	Thr	Asn 330
Ile	Val	Cys	Lys	Leu 335	Asp	Ser	Ser	Gly	Gly 340	Ala	Val	Gln	Leu	
Asp	Thr	Ser	Ile	Ser 350	Ile	His	Val	Pro	Glu 355	Gly	His	Val	Ala	
Gly	Glu	Thr	Gln	Gln 365	Ile	Ser	Met	Lys	Ala 370	Leu	Leu	Asp	Pro	
Leu	Glu	Leu	Asn	Ser 380	qaA	Arg	Ser	Суз	Ser 385	Ile	Ser	Pro	Val	
Glu	Val	Lys	Leu	Ser 395	Asn	Leu	Glu	Val	Lys 400	Thr	Ser	Ile	Ile	
Glu	Met	Lys	Val	Ser 410	Ala	Glu	Ile	Lys	Asn 415	Asp	Leu	Phe	Ser	
Ser	Thr	Val	Gly	Leu 425	Gln	Суѕ	Leu	Arg	Ser 430	Asp	Ser	Lys	Glu	Gly 435
Pro	Tyr	Val	Ser	Val 440	Pro	Leu	Asn	Cys	Ser 445	Суѕ	Gly	Asp	Thr	Val 450
Gln	Ala	Gln	Leu	His 455	Asn	Leu	Glu	Pro	Cys 460	Met	Tyr	Val	Ala	Val 465
			Gly	470					475					480
			Lys	485	54				490		_		_	495
			Ser	500					505		-	_		510
			Lys	515					520					525
			Ala	530					535					540
			Ser	545					550					555
			Asn	560					565					570
			Phe	575					580					585
			Thr	590					595			_		600
			Gln	605					610					615
			Gln	620					625					630
			Gln	635					640					645
			Ser	650					655		_			660
			Pro	665					670					675
			Arg	680					685					690
			Gly	695					700					705
			Leu	710					715					720
			Gly	725					730					735
Arg	Ата	arg	Pro	Ser 740	ren	Суѕ	Ser	Gly	Pro 745	Glu	Leu	Ser	Thr	5er 750

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Val Leu Leu Glu Gln Ile Leu Arg Pro Cys Lys Phe Leu Thr Tyr
                                    760
                755
Ile Tvr Ala Ser Val Arg Thr Leu Leu Met Glu Asn Ile Ser Ser
                770
                                     775
Trp Arg Ser Phe Ala Asp Ala Leu Gly Tyr Val Asn Leu Pro Leu
                                    790
                785
Thr Phe Phe Cys Arg Ala Glu Leu Asp Ser Glu Pro Glu Arg Val
                                    805
                800
Ala Ser Val Leu Glu Lys Leu Lys Glu Asp Cys Asn Asn Thr Glu
                                    820
                815
Asn Lys Glu Arg Lys Ser Phe Gln Lys Glu Leu Val Met Ala Leu
                                    835
                830
Leu Lys Met Asp Cys Gln Gly Leu Val Val Arg Leu Ile Gln Asp
                845
                                    850
Phe Val Leu Leu Thr Thr Ala Val Glu Val Ala Gln Arg Trp Arg
                                     865
                                                         870
                860
Glu Leu Ala Glu Lys Leu Ala Lys Val Ser Lys Gln Gln Met Asp
                                                         885
                                     880
                875
Ala Tyr Glu Ser Pro His Arg Asp Arg Asn Gly Val Val Asp Ser
                890
                                     895
Glu Ala Met Trp Lys Pro Ala Tyr Asp Phe Leu Leu Thr Trp Ser
                905
                                     910
                                                         915
His Gln Ile Gly Asp Ser Tyr Arg Asp Val Ile Gln Glu Leu His
                                     925
                                                         930
                920
Leu Gly Leu Asp Lys Met Lys Asn Pro Ile Thr Lys Arg Trp Lys
                                                         945
                935
                                     940
His Leu Thr Gly Thr Leu Ile Leu Val Asn Ser Leu Asp Val Leu
                                                         960
                950
                                     955
Arg Ala Ala Phe Ser Pro Ala Asp Gln Asp Phe Val Ile
                                     970
                965
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<210> 199
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### <400> 199

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Gly Gly Gly Pro Met Lys Asp Cys Glu Tyr Ser Gln Ile Ser Thr
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His Ser Ser Ser Pro Met Glu Ser Pro His Lys Lys Lys Ile
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                                     25
Ala Ala Arg Arg Lys Trp Glu Val Phe Pro Gly Arg Asn Lys Phe
                 35
Phe Cys Asn Gly Arg Ile Met Met Ala Arg Gln Thr Gly Val Phe
                 50
                                     55
Tyr Leu Thr Leu Val Leu Ile Leu Val Thr Ser Gly Leu Phe Phe
                                     70
                 65
Ala Phe Asp Cys Pro Tyr Leu Ala Val Lys Ile Thr Pro Ala Ile
                                     85
                 80
Pro Ala Val Ala Gly Ile Leu Phe Phe Phe Val Met Gly Thr Leu
                                                         105
                 95
                                    100
Leu Arg Thr Ser Phe Ser Asp Pro Gly Val Leu Pro Arg Ala Thr
                110
                                    115
Pro Asp Glu Ala Ala Asp Leu Glu Arg Gln Ile Asp Ile Ala Asn
                125
                                    130
                                                         135
Gly Thr Ser Ser Gly Gly Tyr Arg Pro Pro Pro Arg Thr Lys Glu
                140
                                    145
Val Ile Ile Asn Gly Gln Thr Val Lys Leu Lys Tyr Cys Phe Thr
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<211> 484

<212> PRT

<213> Homo sapiens

<221> misc_feature

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155
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Cys Lys Ile Phe Arg Pro Pro Arg Ala Ser His Cys Ser Leu Cys
                170
                                     175
Asp Asn Cys Val Glu Arg Phe Asp His His Cys Pro Trp Val Gly
                185
                                     190
Asn Cys Val Gly Lys Arg Asn Tyr Arg Phe Phe Tyr Met Phe Ile
                200
                                     205
Leu Ser Leu Ser Phe Leu Thr Val Phe Ile Phe Ala Phe Val Ile
                215
                                     220
Thr His Val Ile Leu Arg Ser Gln Gln Thr Gly Phe Leu Asn Ala
                230
                                     235
Leu Lys Asp Ser Pro Ala Ser Val Leu Glu Ala Val Val Cys Phe
                245
                                     250
Phe Ser Val Trp Ser Ile Val Gly Leu Ser Gly Phe His Thr Tyr
                260
                                     265
                                                         270
Leu Ile Ser Ser Asn Gln Thr Thr Asn Glu Asp Ile Lys Gly Ser
                275
                                     280
Trp Ser Asn Lys Arg Gly Lys Glu Asn Tyr Asn Pro Tyr Ser Tyr
                290
                                     295
                                                         300
Gly Asn Ile Phe Thr Asn Cys Cys Val Ala Leu Cys Gly Pro Ile
                                     310
                                                         315
Ser Pro Ser Leu Ile Asp Arg Gly Tyr Ile Gln Pro Asp Thr
                320
                                     325
Pro Gln Pro Ala Ala Pro Ser Asn Gly Ile Thr Met Tyr Gly Ala
                335
                                     340
                                                         345
Thr Gln Ser Gln Ser Asp Met Cys Asp Gln Asp Gln Cys Ile Gln
                350
                                     355
Ser Thr Lys Phe Val Leu Gln Ala Ala Ala Thr Pro Leu Leu Gln
                365
                                     370
                                                         375
Ser Glu Pro Ser Leu Thr Ser Asp Glu Leu His Leu Pro Gly Lys
                380
                                     385
                                                         390
Pro Gly Leu Gly Thr Pro Cys Ala Ser Leu Thr Leu Gly Pro Pro
                395
                                     400
Thr Pro Pro Ala Ser Met Pro Asn Leu Ala Glu Ala Thr Leu Ala
                410
                                     415
Asp Val Met Pro Arg Lys Asp Glu His Met Gly His Gln Phe Leu
                425
                                     430
                                                         435
Thr Pro Asp Glu Ala Pro Ser Pro Pro Arg Leu Leu Ala Ala Gly
                440
                                     445
Ser Pro Leu Ala His Ser Arg Thr Met His Val Leu Gly Leu Ala
                455
                                     460
Ser Gln Asp Ser Leu His Glu Asp Ser Val Arg Gly Leu Val Lys
                470
Leu Ser Ser Val
<210> 200
<211> 275
<212> PRT
<213> Homo sapiens
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Gln Arg His Gly His Met Pro Gln Ala Phe Leu Leu Gly Ser Ile
His Glu Pro Ala Gly Ala Leu Met Glu Pro Gln Pro Cys Pro Gly
                 20
                                      25
                                                          30
Ser Leu Ala Glu Ser Phe Leu Glu Glu Glu Leu Arg Leu Asn Ala
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Glu Leu Ser Gln Leu Gln Phe Ser Glu Pro Val Gly Ile Ile Tyr

```
55
Asn Pro Val Glu Tyr Ala Trp Glu Pro His Arg Asn Tyr Val Thr
                  65
Arg Tyr Cys Gln Gly Pro Lys Glu Val Leu Phe Leu Gly Met Asn
                  80
                                      85
                                                           90
Pro Gly Pro Phe Gly Met Ala Gln Thr Gly Val Pro Phe Gly Glu
                  95
                                     100
Val Ser Met Val Arg Asp Trp Leu Gly Ile Val Gly Pro Val Leu
                 110
                                     115
Thr Pro Pro Gln Glu His Pro Lys Arg Pro Val Leu Gly Leu Glu
                 125
                                     130
Cys Pro Gln Ser Glu Val Ser Gly Ala Arg Phe Trp Gly Phe Phe
                 140
                                     145
Arg Asn Leu Cys Gly Gln Pro Glu Val Phe Phe His His Cys Phe
                 155
                                     160
Val His Asn Leu Cys Pro Leu Leu Phe Leu Ala Pro Ser Gly Arg
                 170
                                     175
Asn Leu Thr Pro Ala Glu Leu Pro Ala Lys Gln Arg Glu Gln Leu
                 185
                                     190
Leu Gly Ile Cys Asp Ala Ala Leu Cys Arg Gln Val Gln Leu Leu
                 200
                                     205
                                                          210
Gly Val Arg Leu Val Val Gly Val Gly Arg Leu Ala Glu Gln Arg
                                     220
Ala Arg Arg Ala Leu Ala Gly Leu Met Pro Glu Val Gln Val Glu
                 230
                                     235
                                                         240
Gly Leu Leu His Pro Ser Pro Arg Asn Pro Gln Ala Asn Lys Gly
                 245
                                     250
                                                          255
Trp Glu Ala Val Ala Lys Glu Arg Leu Asn Glu Leu Gly Leu Leu
                260
Pro Leu Leu Lys
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<211> 245
<212> PRT
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Lys Ala Phe Ala Ser Gln Asn Asn Tyr Arg Ile Asp Ala Asn Gln
Glu Leu Leu Ala Ile Gly Leu Thr Asn Met Leu Gly Ser Leu Val
                 20
Ser Ser Tyr Pro Val Thr Gly Ser Phe Gly Arg Thr Ala Val Asn
                 35
                                      40
Ala Gln Ser Gly Val Cys Thr Pro Ala Cly Gly Leu Val Thr Gly
                 50
                                      55
                                                          60
Val Leu Val Leu Ser Leu Asp Tyr Leu Thr Ser Leu Phe Tyr
                 65
                                     70
Tyr Ile Pro Lys Ser Ala Leu Ala Ala Val Ile Ile Met Ala Val
                 80
                                     85
Ala Pro Leu Phe Asp Thr Lys Ile Phe Arg Thr Leu Trp Arg Val
                 95
                                     100
Lys Arg Leu Asp Leu Leu Pro Leu Cys Val Thr Phe Leu Leu Cys
                110
                                    115
Phe Trp Glu Val Gln Tyr Gly Ile Leu Ala Gly Ala Leu Val Ser
                125
                                    130
Leu Leu Met Leu Leu His Ser Ala Ala Arg Pro Glu Thr Lys Val
                140
                                    145
Ser Glu Gly Pro Val Leu Val Leu Gln Pro Ala Ser Gly Leu Ser
```

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Phe Pro Ala Met Glu Ala Leu Arg Glu Glu Ile Leu Ser Arg Ala
                 170
                                     175
 Leu Glu Gly Ala Trp Ala Gly Val Lys Cys Pro Arg His Ala Ala
                 185
                                     190
 Trp Ser Trp Ser Ala Pro Met Ser Ala Ala Ser Thr Thr Leu Trp
                 200
                                     205
 Cys Trp Asp Ser Ala Ser Ser Ser Arg Thr Ser Arg Ser Arg Ala
                 215
                                     220
Ser Pro Trp Pro Leu Trp Ala Cys Arg Ser Pro Phe Ser Val Ser
                 230
                                     235
 Cys Cys Pro Leu Thr
                 245
 <210> 202
 <211> 247
 <212> PRT
 <213> Homo sapiens
 <220>
 <221> misc_feature
 <223> Incyte ID No: LG:233444.9.orf2:2002JAN18
 <400> 202
 Ser Val Ser Cys Leu Val Tyr Met Thr Lys Gly Thr Leu Ala Phe
                                      10
 Ser Asn Ala Trp Thr Tyr Leu Val Ile Ile Asn Asn Met Ser Gln
                                      25
 Leu Phe Ala Met Tyr Cys Leu Leu Phe Tyr Lys Val Leu Lys
                  35
                                      40
 Glu Glu Leu Ser Pro Ile Gln Pro Val Gly Lys Phe Leu Cys Val
                  50
                                      55
 Lys Leu Val Val Phe Val Ser Phe Trp Gln Ala Val Val Ile Ala
                                      70
                  65
 Leu Leu Val Lys Val Gly Val Ile Ser Glu Lys His Thr Trp Glu
                  80
                                      85
 Trp Gln Thr Val Glu Ala Val Ala Thr Gly Leu Gln Asp Phe Ile
                  95
                                     100
 Ile Cys Ile Glu Met Phe Leu Ala Ala Ile Ala His His Tyr Thr
                 110
                                     115
 Phe Ser Tyr Lys Pro Tyr Val Gln Glu Ala Glu Glu Gly Ser Cys
                 125
                                     130
 Phe Asp Ser Phe Leu Ala Met Trp Asp Val Ser Asp Ile Arg Asp
                 140
                                     145
                                                          150
 Asp Ile Ser Glu Gln Val Arg His Val Gly Arg Thr Val Arg Gly
                 155
                                     160
 His Pro Arg Lys Lys Leu Phe Pro Glu Asp Gln Asp Gln Asn Glu
                 170
                                     175
 His Thr Ser Leu Leu Ser Ser Ser Gln Asp Ala Ile Ser Ile
                 185
                                     190
 Ala Ser Ser Met Pro Pro Ser Pro Met Gly His Tyr Gln Gly Phe
                 200
                                     205
 Gly His Thr Val Thr Pro Gln Thr Thr Pro Thr Thr Ala Lys Ile
                 215
                                     220
 Ser Asp Glu Ile Leu Ser Asp Thr Ile Gly Glu Lys Lys Glu Pro
                 230
                                     235
 Ser Asp Lys Ser Val Asp Ser
                 245
 <210> 203
 <211> 749
 <212> PRT
 <213> Homo sapiens
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```
Val Phe Phe Gln Arg Asp Pro Ser Leu Leu Leu His Lys Leu Leu
                                     445
Leu Gly Thr Ser Gly Glu Gly Lys Ala Glu Gly Glu Ser Ser Pro
                455
                                     460
Pro Met Ala Arg Ser Thr Pro Ser Gln Glu Leu Leu Arg Ala Thr
                470
                                     475
Gln Leu His Gln Tyr Val Glu Gly Phe Leu Leu His Gly Leu Leu
                485
                                     490
Pro Ala His Val Ile Arg Leu Leu Lys Pro His Val Gln Ala
                500
                                     505
Gln Gln Asp Leu Gln Leu Leu Glu Leu Glu Lys Met Gly
                515
                                     520
Leu Cys Tyr Cys Leu Asn Lys Pro Lys Gly Lys Pro Leu Asn Gly
                530
                                     535
                                                         540
Ser Thr Ala Trp Tyr Lys Phe Pro Cys Tyr Val Gln Asn Glu Val
                545
                                     550
                                                         555
Pro His Ala Glu Ala Trp Ile Asn Gly Thr Asn Leu Ala Gly Gln
                560
                                     565
Ser Phe Val Ala Glu Gln Leu Gln Ile Glu Tyr Ser Phe Pro Phe
                575
                                     580
                                                         585
Thr Phe Pro Pro Gly Leu Phe Ala Arg Tyr Ser Val Gln Ile Asn
                590
                                     595
Ser His Val Val His Arg Ser Asp Gly Lys Phe Gln Ile Phe Ala
                                     610
Tyr Arg Gly Lys Val Pro Val Val Val Ser Tyr Arg Pro Ala Arg
                620
                                     625
                                                         630
Gly Val Leu Gln Pro Asp Thr Leu Ser Ile Ala Ser His Ala Ser
                635
                                     640
Leu Pro Asn Ile Trp Thr Ala Trp Gln Ala Ile Thr Pro Leu Val
                650
                                     655
Glu Glu Leu Asn Val Leu Leu Gln Glu Trp Pro Gly Leu His Tyr
                665
                                     670
                                                         675
Thr Val His Ile Leu Cys Ser Lys Cys Leu Lys Arg Cly Ser Pro
                680
                                     685
Asn Pro His Ala Phe Pro Gly Glu Leu Leu Ser Gln Pro Arg Pro
                695
                                     700
                                                         705
Glu Gly Val Ala Glu Ile Ile Cys Pro Lys Asn Gly Ser Glu Arg
                710
                                     715
                                                         720
Val Asn Val Ala Leu Val Tyr Pro Pro Thr Pro Thr Val Ile Ser
                725
                                     730
Pro Cys Ser Lys Lys Asn Val Gly Glu Lys His Arg Asn Gln
                740
                                     745
<210> 204
<211> 330
<212> PRT
<213> Homo sapiens
<220>
<221> misc_feature
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Gly Pro Arg Ser Pro Leu Pro Ile Leu Pro Pro Ala Arg Gly Ser
                                      10
Gly His Leu Met Ala Leu Ala Gly Thr Gln Val Gly Pro Pro Pro
                 20
                                      25
Gln Glu Arg Ala Pro Glu Pro Ile Gly Arg Ala Trp Gly Pro Pro
                 35
                                      40
Gly Ile Thr Gln Pro Ser Ala Pro Gly Ala Thr Val Gly Arg Arg
                 50
                                      55
                                                          60
Val Ser Val Ala Ala Gly Pro Trp Leu His Gly Pro His Gly Ser
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Cys Glu Trp Val Arg Leu Pro Gly Ser Gly Asp Arg Gln Arg Thr
                  80
Asp Pro Arg Leu Gly Ser Trp Arg Glu Gly Arg Arg Gly Ala Gly
                 95
                                     100
                                                         105
Gln Pro Gly Ser Asp Thr Val Ser Ser Ser Gly Arg Arg Pro
                110
                                     115
                                                         120
Ala Gly Ser Thr Gln Ala Gly Arg Gly Trp Ala Ser Leu Glu Pro
                125
                                     130
Ala Thr Ala Leu Val Gly Thr Trp Arg Arg Ala His Val Ser Pro
                140
                                     145
                                                         150
His Ala Ser His Arg Gly Ala Leu Ala Arg Arg Pro Ala Arg Gly
                155
                                     160
Ala Cys Ala Trp Asp Gly Ser Gln Asn Gln Arg Ala Pro Val Arg
                170
                                     175
                                                         180
Leu Ala Ser Thr Val Gly Leu Trp Glu Ser Leu Leu Phe Ile Phe
                185
                                     190
                                                         195
Lys His Leu Gly Phe Ser Thr Gly Ser Trp Leu Leu Phe Pro Gln
                200
                                     205
                                                         210
Gly Met Ser Leu Arg Ser Arg Thr Arg Trp Gly Ser Gln Glu Ala
                215
                                     220
                                                         225
Ala Ala Gln Ser Leu His Ala Gly Lys Gly Ser His Leu Ser Gly
                230
                                    235
                                                         240
Val Gly Ser Leu Val Val Gln Gly Ser Ala Gly Gln Ser Leu Gly
                245
                                    250
                                                         255
Cys Ala Ile Thr Ala Thr Ala Phe Leu Leu Gly Ala Ser Thr Ser
                260
                                     265
                                                         270
His Pro Lys Thr Gly Pro Cys Ala Ser Pro Thr Arg Gly Glu Arg
                275
                                    280
Thr Arg Pro Arg Arg Gln Gly Pro Glu Tyr Leu Gly Gly Gly Asp
                290
                                    295
Thr Pro Arg Gly His Arg Gly Gly Ser His Leu Gly Thr Cys Leu
                305
                                    310
Leu Pro Phe Ser Ser Ser Thr Cys Gln Val Gly Pro Trp Val
                320
                                    325
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<213> Homo sapiens

<220>

<221> misc_feature

<223> Incyte ID No: LG:236649.14.orf1:2002JAN18

<400> 205

Arg Ala Trp Gly Pro Leu Leu Ser Ile Gly Gly Gln Arg Gln His Gln Arg Leu Val Glu Gln Tyr Arg Clu Gln Ser Trp Met Thr Met Ala Asn Leu Glu Lys Glu Leu Gln Glu Met Glu Ala Arg Tyr Glu Lys Glu Phe Gly Asp Gly Ser Asp Glu Asn Glu Met Glu Glu His Glu Leu Lys Asp Glu Glu Asp Gly Lys Asp Ser Asp Glu Ala Glu Asp Ala Glu Leu Tyr Asp Asp Leu Tyr Cys Pro Ala Cys Asp Lys Ser Phe Lys Thr Glu Lys Ala Met Lys Asn His Glu Lys Ser Lys Lys His Arg Glu Met Val Ala Leu Leu Lys Gln Gln Leu Glu Glu Glu Glu Asn Phe Ser Arg Pro Gln Ile Asp Glu Asn Pro

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125
                                    130
                                                         135
Leu Asp Asp Asn Ser Glu Glu Glu Met Glu Asp Ala Pro Lys Gln
                140
                                    145
Lys Leu Ser Lys Lys Gln Lys Lys Lys Gln Lys Pro Ala Gln
                155
                                    160
Asn Tyr Asp Asp Asn Phe Asn Val Asn Gly Pro Gly Glu Gly Val
                170
                                    175
Lys Val Asp Pro Glu Asp Thr Asn Leu Asn Gln Asp Ser Ala Lys
                185
                                    190
Glu Leu Glu Asp Ser Pro Gln Glu Asn Val Ser Val Thr Glu Ile
                200
                                    205
Ile Lys Pro Cys Asp Asp Pro Lys Ser Glu Ala Lys Ser Val Pro
                215
                                    220
Lys Pro Lys Gly Lys Lys Thr Lys Asp Met Lys Lys Pro Val Arg
                230
                                    235
Val Pro Ala Glu Pro Gln Thr Met Ser Val Leu Ile Ser Cys Thr
                245
                                    250
Thr Cys His Ser Glu Phe Pro Ser Arg Asn Lys Leu Phe Asp His
                260
                                    265
                                                         270
Leu Lys Ala Thr Gly His Ala Arg Ala Pro Ser Ser Ser Leu
                275
                                    280
Asn Ser Ala Thr Ser Ser Gln Ser Lys Lys Glu Lys Arg Lys Asn
                290
                                    295
Arg
<210> 206
<211> 213
<212> PRT
<213> Homo sapiens
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Ala Thr Asp Thr Ser Asp Thr Arg Cys Gly Gly Arg Arg Ser Gly
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Ser Gly Ala Val Pro Ser Ala Ser Val Gly Arg His Pro Leu Ala
                 20
                                     25
Ala Glu Pro Ala Leu His Trp Gly Pro Trp Gly Arg Leu Leu Gln
                 35
                                     40
Thr Ser Leu Arg Val Val Gly Cys Cys Arg Val Gly Leu Gly Asp
                 50
                                     55
Ala Thr Gly Ser Gly Ser Gly Ala Ala Gly Pro Ser Ser His
Cys Pro Ser Ser Asp Pro Thr Val Leu Trp Lys Leu Val Gln Gly
                 80
                                     85
Thr Cys His Cys Asp His Leu Asp Ala Asp Thr Cys Phe Pro Thr
                 95
                                    100
Thr Ala Arg Lys Asn His Gly Pro Gly Ser Leu Ser Phe Gly Asp
                110
                                    115
Val Ala Val Gly Phe Thr Arg Lys Glu Trp Leu Ala Ala Gly Pro
                125
                                    130
Gly Ala Glu Asp Pro Val Pro Gly Cys Asn Ala Gly Glu Leu Gln
                140
                                    145
                                                         150
Pro Pro Ala Leu Cys Gly Leu Asn Cys Leu Leu Ala Trp Lys Ala
                155
                                    160
Gln Leu Ala Gln His Lys Ser Phe Asn Arg Phe Ser Pro Arg Gly
                170
                                    175
Cys Gln Val Ser Lys Pro Ala Val Ile Ser Ser Leu Glu Gln Gly
                185
                                    190
Lys Glu Pro Trp Met Glu Glu Glu Glu Ile Arg Thr Trp Ser Phe
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200 205 210 Pro Glu Ser <210> 207 <211> 322 <212> PRT <213> Homo sapiens <220>. <221> misc_feature <223> Incyte ID No: LG:335727.8.orf2:2002JAN18 <400> 207 Lys Ala Gly Gly Thr Asn Cys Gln Cys Leu Glu Ser Gly Ser Arg 10 Leu Pro Phe Leu Gly Ala Leu Gly Val Ala Val Ala Glu 20 25 Ile Gly Cys Thr Met Ser Ala Phe Glu Lys Pro Gln Ile Ile Ala 35 40 His Ile Gln Lys Gly Phe Asn Tyr Thr Val Phe Asp Cys Lys Trp 50 55 Val Pro Cys Ser Ala Lys Phe Val Thr Met Gly Asn Phe Ala Arg 65 Gly Thr Gly Val Ile Gln Leu Tyr Glu Ile Gln His Gly Asp Leu 80 85 Lys Leu Leu Arg Glu Ile Glu Lys Ala Lys Pro Ile Lys Cys Gly 95 100 105 Thr Phe Gly Ala Thr Ser Leu Gln Gln Arg Tyr Leu Ala Thr Gly 110 115 Asp Phe Gly Gly Asn Leu His Ile Trp Asn Leu Glu Ala Pro Glu 125 130 Met Pro Val Tyr Ser Val Lys Gly His Lys Glu Ile Ile Asn Ala 140 145 Ile Asp Gly Ile Gly Gly Leu Gly Ile Gly Glu Gly Ala Pro Glu 155 160 Ile Val Thr Gly Ser Arg Asp Gly Thr Val Lys Val Trp Asp Pro 170 175 180 Arg Gln Lys Asp Asp Pro Val Ala Asn Met Glu Pro Val Gln Gly 185 190 195 Glu Asn Lys Arg Asp Cys Trp Thr Val Ala Phe Gly Asn Ala Tyr 200 205 Asn Gln Glu Glu Arg Val Val Cys Ala Gly Tyr Asp Asn Gly Asp 215 220 Ile Lys Leu Phe Asp Leu Arg Asn Met Ala Leu Arg Trp Glu Thr 230 235 Asn Ile Lys Asn Gly Val Cys Ser Leu Glu Phe Asp Arg Lys Asp 245 250 Ile Ser Met Asn Lys Leu Val Ala Thr Ser Leu Glu Gly Lys Phe 260 265 His Val Phe Asp Met Arg Thr Gln His Pro Thr Lys Gly Phe Ala 275 280 Ser Val Ser Glu Lys Ala His Lys Ser Thr Val Trp Gln Val Arg 290 295 His Leu Pro Gln Asn Arg Glu Leu Phe Leu Thr Ala Gly Gly Ala 305 310 Gly Gly Leu His Leu Trp Lys 320 <210> 208 <211> 238 <212> PRT <213> Homo sapiens

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<221> misc feature
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Ala Val Ser Leu Arg Cys Leu Gly Tyr Lys Leu Gly Ser Leu Leu
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Thr Thr Asn His Ser Phe Phe Phe Phe His Leu Gln Pro Leu Thr
                                     25
Thr Asp Leu His Tyr Tyr Tyr Ile Leu Glu Leu Ser Phe Tyr Trp
                 35
                                     40
Ser Leu Met Phe Ser Gln Phe Thr Asp Ile Lys Arg Lys Asp Phe
                 50
                                     55
Gly Ile Met Phe Leu His His Leu Val Ser Ile Phe Leu Ile Thr
                                     70
Phe Ser Tyr Val Asn Asn Met Ala Arg Val Gly Thr Leu Val Leu
                 80
                                     85
Cys Leu His Asp Ser Ala Asp Ala Leu Leu Glu Ala Ala Lys Met
                95
                                    100
Ala Asn Tyr Ala Lys Phe Gln Lys Met Cys Asp Leu Leu Phe Val
                110
                                    115
Met Phe Ala Val Val Phe Ile Thr Thr Arg Leu Gly Ile Phe Pro
                125
                                   130
Leu Trp Val Leu Asn Thr Thr Leu Phe Glu Ser Trp Glu Ile Val
                140
                                    145
Gly Pro Tyr Pro Ser Trp Trp Val Phe Asn Leu Leu Leu Leu
                155
                                    160
Val Gln Gly Leu Asn Cys Phe Trp Ser Tyr Leu Ile Val Lys Ile
                170
                                    175
Ala Cys Lys Ala Val Ser Arg Gly Lys Ala Gly Lys Trp Asn Pro
                185
                                   · 190
Tyr Met Cys Pro Arg Met Ile Glu Val Ile Leu Ser Leu Ala Gln
                200
                                    205
Met Arg Arg Thr Gln Asn Leu Arg Glu Arg Ile Pro Thr Leu Arg
                215
                                    220
Gln Pro Pro Met Gly Pro Val Val Pro Thr Gly Ile Ser
                230
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For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: MOLECULES FOR DISEASE DETECTION AND TREATMENT

(57) Abstract: The present invention provides purified disease detection and treatment molecule polynucleotides (mddt). Also encompassed are the polypeptides (MDDT) encoded by mddt. The invention also provides for the use of mddt, or complements, oligonucleotides, or fragments thereof in diagnostic assays. The invention further provides for vectors and host cells containing mddt for the expression of MDDT. The invention additionally provides for the use of isolated and purified MDDT to induce antibodies and to screen libraries of compounds and the use of anti-MDDT antibodies in diagnostic assays. Also provided are microarrays containing mddt and methods of use.



# INTERNATIONAL SEARCH REPORT

International application No.

PCT/US03/01363

A. CLAS IPC(7)	SSIFICATION OF SUBJECT MATTER						
US CL	: C12N 15/11, 15/12, 15/00; C12Q 1/68; C07K : 536/23.1, 23.5; 435/6, 320/1, 325, 252.3; 530	7/350- 514/12					
According to	International Patent Classification (IPC) or to both n	national classification and IPC					
	DS SEARCHED						
Minimum do U.S.: 53	cumentation searched (classification system followed 36/23.1, 23.5; 435/6, 320/1, 325, 252.3; 530/350; 5	by classification symbols) 14/12					
Documentation	on searched other than minimum documentation to the	e extent that such documents are included in	n the fields searched				
Electronic da Compugen, S	ta base consulted during the international search (nan EEQ ID NOs: 1 and 105	ne of data base and, where practicable, sear	ch terms used)				
	UMENTS CONSIDERED TO BE RELEVANT						
Category *	Citation of document, with indication, where	appropriate, of the relevant passages	Relevant to claim No.				
. X	WO 01/53312 A (HYSEQ, INC.) 26 July 2001 (26	5.07.2001), compare SEQ ID NO: !,	3 and 6-8				
Y	position 529-1305 to reference SEQ ID NO: 1215, attached to the reference.	positions 403-1179 and see alignment	16 and 20-26				
			10 and 20-20				
x	EP 1033401 A2 (GENSET) 06 September 2000 (06 positions 25-140 to SEQ ID NO: 7103, positions 1-	.09.2000), compare SEQ ID NO: 105, 116 and alignment attached to reference.	27				
	documents are listed in the continuation of Box C.	See patent family annex.					
	ecial categories of cited documents:	"T" later document published after the inter date and not in conflict with the applica	national filing date or priority				
"A" document of particular	defining the general state of the art which is not considered to be ar relevance	principle or theory underlying the inver	ition				
"E" carlier app	lication or patent published on or after the international filing date	"X" document of particular relevance; the considered novel or cannot be considered	aimed invention cannot be ad to involve an inventive step				
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"O" document	referring to an oral disclosure, use, exhibition or other means	combined with one or more other such of being obvious to a person skilled in the	documents, such combination				
"P" document priority date	published prior to the international filing date but later than the te claimed	"&" document member of the same patent family					
Date of the ac	tual completion of the international search	Date of mailing of the international search	report				
26 August 200	3 (26.08.2003)	UD DEC 2003 A					
rvanie and mai Mail	iling address of the ISA/US Stop PCT, Attn: ISA/US	Authorized officer					
Com	missioner for Patents	James Martinell					
	Box 1450 andria, Virginia 22313-1450	Telephone No. (703) 308-0196	Ille Illian.				
	/210 (second sheet) (July 1998)	<u> </u>	2				
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BNSDOCID: <WO_____03062379A3_I_>

# INTERNATIONAL SEARCH REPORT

International application No.

PCT/US03/01363

Box I Observations where certain claims were found unsearchable (Continuation of Item 1 of first sheet)					
This international report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:					
1. Claim Nos.: because they relate to subject matter not required to be searched by this Authority, namely:					
2. Claim Nos.:  because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:					
3. Claim Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).					
Box II Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)					
This International Searching Authority found multiple inventions in this international application, as follows:					
-					
As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.					
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.					
As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.: Please See Continuation Sheet					
4. No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:					
Remark on Protest					
No protest accompanied the payment of additional search fees.					
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Form PCT/ISA/210 (continuation of first sheet(1)) (July 1998)

BNSDOCID: <WO____03062379A3_l_>

# PCT/US03/01363 INTERNATIONAL SEARCH REPORT Continuation of Box II Item 3: 1-10, 12, 13, 16, 17, and 19-28 to the extent that they include SEQ ID NO: 1 and 105

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